

GenCore version 5.1.4\_p5\_4578  
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protein search, using sw model

May 16, 2003, 10:12:16 ; Search time 35 Seconds

167.515 Million cell updates/sec

SEQ1-4EDITS  
197

1 ANSFLXXLRqgSLXRCIXX.....XXAKXIFedVDDTLAFWSKH 44

BLOSUM62	
Gapon 10 0	Gapext 0 5

908470 seqs, 133250620 residues

hits satisfying chosen parameters: 908470

Length: 20000000000

Minimum Match 08

Listing first 45 summaries

A\_Geneseq\_101002:\*

1:	/SIDS2/gcgdata/geneseq/geneseqp-emb1/AA1981.DAT *
2:	/SIDS2/gcgdata/geneseq/geneseqp-emb1/AA1981.DAT *
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4:	/SIDS2/gcgdata/geneseq/geneseqp-emb1/AA1983.DAT *
5:	/SIDS2/gcgdata/geneseq/geneseqp-emb1/AA1984.DAT *
6:	/SIDS2/gcgdata/geneseq/geneseqp-emb1/AA1985.DAT *
7:	/SIDS2/gcgdata/geneseq/geneseqp-emb1/AA1986.DAT *
8:	/SIDS2/gcgdata/geneseq/geneseqp-emb1/AA1987.DAT *
9:	/SIDS2/gcgdata/geneseq/geneseqp-emb1/AA1988.DAT *
10:	/SIDS2/gcgdata/geneseq/geneseqp-emb1/AA1989.DAT *
11:	/SIDS2/gcgdata/geneseq/geneseqp-emb1/AA1990.DAT *
12:	/SIDS2/gcgdata/geneseq/geneseqp-emb1/AA1992.DAT *
13:	/SIDS2/gcgdata/geneseq/geneseqp-emb1/AA1993.DAT *
14:	/SIDS2/gcgdata/geneseq/geneseqp-emb1/AA1994.DAT *
15:	/SIDS2/gcgdata/geneseq/geneseqp-emb1/AA1994.DAT *
16:	/SIDS2/gcgdata/geneseq/geneseqp-emb1/AA1995.DAT *
17:	/SIDS2/gcgdata/geneseq/geneseqp-emb1/AA1996.DAT *
18:	/SIDS2/gcgdata/geneseq/geneseqp-emb1/AA1997.DAT *
19:	/SIDS2/gcgdata/geneseq/geneseqp-emb1/AA1998.DAT *
20:	/SIDS2/gcgdata/geneseq/geneseqp-emb1/AA1999.DAT *
21:	/SIDS2/gcgdata/geneseq/geneseqp-emb1/AA2000.DAT *
22:	/SIDS2/gcgdata/geneseq/geneseqp-emb1/AA2001.DAT *
23:	/SIDS2/gcgdata/geneseq/geneseqp-emb1/AA2002.DAT *

Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed and is derived by analysis of the total score distribution.

## SUMMARIES

Result No.	Score	Query Match	length	DB	ID	Description
1	179	50.9	44	20	AAV18300	Modified GLA domain
2	179	50.9	419	22	AAE08630	Human protein C de
3	179	50.9	419	22	AAB82677	Human protein C de
4	179	50.9	419	22	AAB82678	Human protein C de
5	176	89.3	44	20	AAV18301	Modified GLA domain
6	174	88.3	419	22	AAE08627	Human protein C de
7	174	88.3	419	22	AAE08628	Human protein C de
8	174	88.3	419	22	AAE08629	Human protein C de
9	174	88.3	419	22	AAB82675	Human protein C de
10	174	88.3	419	22	AAB82676	Human protein C de

11	173	87.8	44	20	AAV18298	Modified GLA domain
12	170	86.3	44	20	AAV18299	Modified GLA domain
13	168	85.3	44	20	AAV18307	Modified GLA domain
14	168	85.3	44	20	AAV18297	Modified GLA domain
15	160	81.2	44	20	AAV18309	Modified GLA domain
16	160	81.2	44	20	AAV18303	Human protein C GLA
17	160	81.2	44	22	AAV83640	Human protein C
18	160	81.2	45	19	AAV75710	Partial human protein C
19	160	81.2	415	21	AAV56803	Truncated human protein C (PC). H
20	160	81.2	419	14	AAV85760	Primary structure
21	160	81.2	419	19	AAV82753	Human mature wild
22	160	81.2	419	22	AAV86525	Wild-type human protein C
23	160	81.2	419	22	AAV82673	Human protein C de
24	160	81.2	419	22	AAV86894	Human protein C de
25	160	81.2	419	22	AAV86896	Human protein C de
26	160	81.2	419	22	AAV86897	Human protein C de
27	160	81.2	419	22	AAV86898	Human protein C de
28	160	81.2	419	23	AAV99002	Human protein C
29	160	81.2	419	23	AAV99003	Human protein C
30	160	81.2	419	23	AAV99004	Human protein C
31	160	81.2	419	23	AAV99005	Human protein C
32	160	81.2	419	23	AAV99006	Human protein C
33	160	81.2	419	23	AAV99007	Human protein C
34	160	81.2	419	23	AAV99008	Human protein C
35	160	81.2	419	23	AAV99009	Human protein C
36	160	81.2	419	23	AAV99010	Human protein C
37	160	81.2	419	23	AAV99011	Human protein C
38	160	81.2	419	23	AAV99012	Human protein C
39	160	81.2	419	23	AAV99013	Human protein C
40	160	81.2	419	23	AAV99014	Human protein C
41	160	81.2	419	23	AAV99015	Human protein C
42	160	81.2	419	23	AAV99016	Human protein C
43	160	81.2	419	23	AAV99017	Human protein C
44	160	81.2	419	23	AAV99018	Human protein C
45	160	81.2	419	23	AAV99019	Human protein C

## ALIGNMENTS

RESULT 1	
AAY18300	
ID	AAY18300 standard; peptide; 44 AA.
XX	
AC	AAY18300;
XX	
DT	17-AUG-1999 (first entry)
XX	
DE	Modified GLA domain of vitamin K-dependent protein.
KW	GLA domain; muten; vitamin K-dependent protein; clotting disorder;
KW	therapy.
XX	
OS	Homo sapiens.
OS	Synthetic.
XX	
EH	key
FT	Misc-difference 1..44
FT	/note= "Xaa= gamma-carboxyglutamic acid, or glutamic acid"
XX	
PN	MO9920767-Al.
XX	
PD	29-APR-1999.
XX	
PE	20-OCT-1998; 98WO-US22152.
-XX	
PR	23-OCT-1997; 97US-0955636.
XX	
PA	(MINU ) UNIV MINNESOTA.
XX	
PI	Nelstøtuen GL;
XX	

DR WPI: 1999-288309/24.  
 XX  
 XX Vitamin K-dependent polypeptide with modified gamma-carboxyglutamic  
 PT acid domain, useful for treating clotting disorders  
 XX  
 XX Claim 9; Page 79; 86pp; English.  
 PS  
 CC This sequence represents a modified GUA (gamma-carboxyglutamic acid)  
 CC domain. The invention relates to a vitamin K-dependent polypeptide  
 CC comprising a modified GUA domain containing an amino acid substitution  
 CC which enhances membrane binding of the modified polypeptide as compared  
 CC to the native polypeptide. The polypeptide is used to treat a clotting  
 CC disorder by decreasing or increasing clot formation. Modification of the  
 CC GUA domain results in a protein which has enhanced membrane binding  
 CC affinity as compared to the native protein.  
 XX  
 SQ Sequence 44 AA;  
 Query Match 90.9%; Score 179; DB 20; Length 44;  
 Best Local Similarity 100.0%; Pred. No. 1.3e-22;  
 Matches 44; Conservative 0; Mismatches 0; Indels 0; Gaps 0;  
 OY 1 ANSFLXXLRQGSIXRCIXXICDFXXAKXIFEDVDTLAFWSKH 44  
 DB 1 ANSFLXXLRQGSIXRCIXXICDFXXAKXIFEDVDTLAFWSKH 44  
 RESULT 2  
 AAE08630 standard; Protein: 419 AA.  
 AC AAE08630;  
 XX  
 DT 01-NOV-2001 (first entry)  
 XX  
 DE Human protein C derivative #4.  
 XX  
 KW Human; protein C derivative; anticoagulation activity; thrombosis;  
 KW serpin inactivation; acute coronary syndrome; myocardial infarction;  
 KW vascular occlusive disorder; hypercoagulable state; angina; sepsis;  
 KW disseminated intravascular coagulation; DIC; burn; transplantation;  
 KW sickle cell disease; viral haemorrhagic fever; protein C deficiency;  
 KW haemolytic uremic syndrome; acute arterial thrombotic occlusion;  
 KW thromboembolism; prothrombotic disorder; gene therapy; thalassaemia.  
 XX  
 OS Homo sapiens.  
 XX  
 PN WO200159084-A1.  
 XX  
 PD 16-AUG-2001.  
 XX  
 PF 02-FEB-2001; 2001WO-US01221.  
 XX  
 PR 11-FEB-2000; 2000US-0181948.  
 PR 14-MAR-2000; 2000US-0189199.  
 XX  
 PA (ELIT ) LILLY & CO ELI.  
 XX  
 PI Gerlitz BE, Grinnell BW, Jones BE;  
 XX  
 DR WPI: 2001-514662/56.  
 DR N-PSDB; AAD15228.  
 XX  
 PT Protein C derivative for treating acute coronary syndromes, vascular  
 PT occlusive disorders, thrombotic disorders and sepsis, comprises  
 PT substitutions at specified amino acid positions  
 XX  
 PS Claim 6; Page 50-51; 59pp; English.  
 XX  
 CC The invention relates to human protein C derivatives and nucleic acid  
 CC molecules encoding such derivatives. These derivatives have increased  
 CC anticoagulation activity, resistance to serpin inactivation and  
 CC increased sensitivity to thrombin activation compared to wild type

CC protein C, and retains the biological activity of the wild type human  
 CC protein C. Protein C derivatives are useful in the manufacture of a  
 CC medicament for the treatment of acute coronary syndromes e.g. myocardial  
 CC infarction and unstable angina; and disease states predisposing to  
 CC thrombosis; vascular occlusive disorders and hypercoagulable states e.g.  
 CC disseminated intravascular coagulation (DIC), burns, transplantations,  
 CC thalassaemia, sickle cell disease, viral haemorrhagic fever and  
 CC haemolytic uremic syndrome; sepsis in combination with bacterial  
 CC permeability increasing protein; thrombotic disorders in combination  
 CC with an anti-platelet agent; protein C deficiency; acute arterial  
 CC thrombotic occlusion, thromboembolism or stenosis in coronary, cerebral  
 CC or peripheral arteries or in vascular grafts in combination with a  
 CC thrombolytic agent. Nucleic acid molecules of the invention are useful  
 CC for treating humans with genetically predisposed prothrombotic disorders  
 CC by gene therapy. The present sequence is human protein C derivative.  
 XX  
 SQ Sequence 419 AA;  
 Query Match 90.9%; Score 179; DB 22; Length 419;  
 Best Local Similarity 79.5%; Pred. No. 1.4e-21;  
 Matches 35; Conservative 0; Mismatches 9; Indels 0; Gaps 0;  
 OY 1 ANSFLXXLRQGSIXRCIXXICDFXXAKXIFEDVDTLAFWSKH 44  
 DB 1 ANSFLXXLRQGSIXRCIXXICDFXXAKXIFEDVDTLAFWSKH 44  
 RESULT 3  
 AAB82677 standard; Protein: 419 AA.  
 AC AAB82677;  
 XX  
 DT 15-OCT-2001 (first entry).  
 XX  
 DE Human protein C derivative (H10Q/S11C/Q32E/N33D/L194S).  
 XX  
 KW Protein C; human; coronary syndrome; thrombosis; angina;  
 KW myocardial infarction; vascular occlusive disorder;  
 KW hypercoagulable; sepsis; protein C deficiency; occlusion;  
 KW thromboembolism; stenosis; antibacterial; immunosuppressive;  
 KW thrombolytic; caditant; antianginal; anticoagulant; therapy;  
 KW mutant; mutlein.  
 XX  
 OS Homo sapiens.  
 XX  
 PN Synthetic.  
 XX  
 FH Key Location/Qualifiers  
 FT Misc-difference 10  
 FT Misc-difference 11  
 FT Misc-difference 11  
 FT Misc-difference 11  
 FT Misc-difference 32  
 FT Misc-difference 33  
 FT Misc-difference 33  
 FT Misc-difference 33  
 FT Misc-difference 33  
 FT Domain  
 FT Disulfide-bond 50..69  
 FT Disulfide-bond 59..64  
 FT Disulfide-bond 80..89  
 FT Disulfide-bond 98..109  
 FT Disulfide-bond 120..133  
 FT Disulfide-bond 141..217  
 FT Disulfide-bond 196..212  
 FT Disulfide-bond 331..345  
 FT Disulfide-bond 356..384  
 FT Cleavage-site 156..157  
 FT /note= "Gla domain"  
 FT /note= "Gln in wild-type protein"  
 FT /note= "Asn in wild-type protein"  
 FT /note= "Leu in wild-type protein"  
 FT /note= "Gla domain"  
 FT /note= "cleavage makes a 2-chain inactive  
 FT precursor (155-amino acid light chain  
 FT attached via a disulfide bond to a



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FT Modified-site 20 /note= "gamma-carboxylated"
FT Modified-site 25 /note= "gamma-carboxylated"
FT Modified-site 26 /note= "gamma-carboxylated"
FT Peptide 158..169 /note= "gamma-carboxylated"
FT 158..169 /note= "activation peptide; removal activates the
FT 2-chain zymogen"
FT Cleavage-site 169..170 /note= "thrombin cleavage site"
FT Modified-site 29 /note= "N-glycosylated"
FT Modified-site 248 /note= "N-glycosylated"
FT Modified-site 313 /note= "N-glycosylated"
FT Modified-site 329 /note= "N-glycosylated"
FT Modified-site /note= "N-glycosylated"
PN WO200157193-A2.
XX
XX
XX 09-AUG-2001.
PD
XX
XX 19-JAN-2001; 2001WO-US00020.
PE
XX
XX 02-FEB-2000; 2000US-0179801.
PR 14-MAR-2000; 2000US-0189197.
XX
XX (ELIL ) LILLY & CO ELI.
XX
XX Gerlitz BE, Jones BE;
XX WPI; 2001-496919/54.
XX
XX Novel human protein C derivative for treating, e.g., myocardial
PT infarction, unstable angina, sepsis, thrombotic disorders, acute
PT arterial thrombotic occlusion, and thromboembolism -
XX
XX Claim 6; Page 56-57; 63pp; English.
XX
XX The present sequence is that of a claimed human protein C derivative
CC in which His at position 10 of the wild-type protein C sequence (see
CC AA82673) is substituted with Gln, Ser at position 11 with Gly, Gln
CC at position 32 with Gln, Asn at position 33 with Asp, Leu at position
CC 194 with Ser, and Thr at position 254 with Ser. It is an example of
CC protein C derivatives of the invention that have at least 2 amino acid
CC substitutions, but which have increased anticoagulant activity and
CC resistance to inactivation by serpins compared with the wild-type
CC protein, while retaining the biological activity of the wild-type
CC protein. A method of producing the derivatives using recombinant
CC DNA methods is claimed. The protein C derivatives are useful for
CC treating coronary syndromes and disease states predisposing to
CC thrombosis (e.g., myocardial infarction and unstable angina).
CC vascular occlusive disorders and hypercoagulable states, sepsis (in
CC combination with bactericidal permeability increasing protein or
CC with tissue factor pathway inhibitor), thrombotic disorders (in
CC combination with an anti-platelet agent or by local delivery through
CC an intracoronary catheter), protein C deficiency, acute arterial
CC thrombotic occlusion, thromboembolism, or stenosis in coronary,
CC cerebral or peripheral arteries or in vascular grafts. Human
CC patients with genetically predisposed prothrombotic disorders may
CC be treated by gene therapy (all claimed).
XX
XX Sequence 419 AA;
SQ

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Query Match 90.9%; Score 179; DB 22; Length 419;  
 Best Local Similarity 79.5%; Pred. No. 1.4e-21;  
 Matches 35; Conservative 0; Mismatches 9; Indels 0; Gaps 0;

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OY 1 ANSFLLXLRQSLXRXCIIXICDFXXAKXIFEDVDITAFMSKH 44
DB 1 ANSFLELRQSLRRECIIEICDFEAKEIFEDVDITAFMSKH 44

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RESULT 5
AAV18301
ID AAV18301 standard; peptide; 44 AA.
XX
XX AAV18301;
AC
AC 17-AUG-1999 (first entry)
DT
XX
XX Modified GLA domain of vitamin K-dependent protein.
DE
XX GLA domain; mutein; vitamin K-dependent protein; clotting disorder;
XX therapy.
KW
KW Homo sapiens.
OS
OS Synthetic.
XX
XX Key Location/Qualifiers
FH Misc-difference 1..44 /note= "Xaa= gamma-carboxyglutamic acid, or glutamic
FT acid"
FT
FT
PN WO9920767-A1.
XX
XX 29-APR-1999.
PD
XX
XX 20-OCT-1998; 98WO-US22152.
XX
XX 23-OCT-1997; 97US-0955636.
XX
XX (MIND ) UNIV MINNESOTA.
XX
XX Nelstuen GL;
XX
XX WPI; 1999-288309/24.
XX
XX Vitamin K-dependent polypeptide with modified gamma-carboxyglutamic
PT acid domain, useful for treating clotting disorders
PT
XX
XX Claim 9; Page 82; 86pp; English.
XX
XX This sequence represents a modified GLA (gamma-carboxyglutamic acid)
CC domain. The invention relates to a vitamin K-dependent polypeptide
CC comprising a modified GLA domain containing an amino acid substitution
CC which enhances membrane binding of the modified polypeptide as compared
CC to the native polypeptide. The polypeptide is used to treat a clotting
CC disorder by decreasing or increasing clot formation. Modification of the
CC GLA domain results in a protein which has enhanced membrane binding
CC affinity as compared to the native protein.
XX
XX Sequence 44 AA;
SQ

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Query Match 89.3%; Score 176; DB 20; Length 44;  
 Best Local Similarity 97.7%; Pred. No. 4.1e-22;  
 Matches 43; Conservative 1; Mismatches 0; Indels 0; Gaps 0;

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OY 1 ANSFLLXLRQSLXRXCIIXICDFXXAKXIFEDVDITAFMSKH 44
DB 1 ANSFLELRQSLRRECIIEICDFEAKEIFEDVDITAFMSKH 44

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RESULT 6  
 AAEO8627  
 ID AAEO8627 standard; Protein; 419 AA.  
 XX  
 XX AAEO8627;  
 AC  
 AC 01-NOV-2001 (first entry)  
 DT  
 XX Human protein C derivative #1.  
 XX  
 XX Human; protein C derivative; anticoagulation activity; thrombosis;  
 KW

KM serpin inactivation; acute coronary syndrome; myocardial infarction;  
 KM vascular occlusive disorder; hypercoagulable state; angina; sepsis;  
 KM disseminated intravascular coagulation; DIC; burn; transplantation;  
 KM sickle cell disease; viral haemorrhagic fever; protein C deficiency;  
 KM haemolytic uremic syndrome; acute arterial thrombotic occlusion;  
 KM thrombocytopenia; prothrombotic disorder; gene therapy; thalassemia.  
 OS Homo sapiens.  
 XX  
 XX WO200159084-A1.  
 PN 16-AUG-2001.  
 PD  
 XX  
 XX 02-FEB-2001; 2001WO-US01221.  
 PF  
 XX 11-FEB-2000; 2000US-0181948.  
 PR 14-MAR-2000; 2000US-0189199.  
 XX  
 XX (ELIL ) LILLY & CO ELI.  
 PA  
 XX  
 XX Gerlitz BE, Grinnell BW, Jones BE;  
 PI  
 XX WPI: 2001-514662/56.  
 DR N-PSDB: AAD15225.  
 XX  
 XX Protein C derivative for treating acute coronary syndromes, vascular  
 PT occlusive disorders, thrombotic disorders and sepsis, comprises  
 PT substitutions at specified amino acid positions  
 PS  
 XX Claim 3; Page 46-47; 59pp; English.  
 XX  
 XX The invention relates to human protein C derivatives and nucleic acid  
 CC molecules encoding such derivatives. These derivatives have increased  
 CC anticoagulation activity, resistance to serpin inactivation and  
 CC increased sensitivity to thrombin activation compared to wild type  
 CC protein C, and retains the biological activity of the wild type human  
 CC protein C. Protein C derivatives are useful in the manufacture of a  
 CC medicament for the treatment of acute coronary syndromes e.g. myocardial  
 CC infarction and unstable angina; and disease states predisposing to  
 CC thrombosis; vascular occlusive disorders and hypercoagulable states e.g.  
 CC disseminated intravascular coagulation (DIC), burns, transplantations,  
 CC thalassemia, sickle cell disease, viral haemorrhagic fever and  
 CC haemolytic uremic syndrome; sepsis in combination with bacterial  
 CC permeability increasing protein; thrombotic disorders in combination  
 CC with an anti-platelet agent; protein C deficiency; acute arterial  
 CC thrombotic occlusion, thromboembolism or stenosis in coronary, cerebral  
 CC or peripheral arteries or in vascular grafts in combination with a  
 CC thrombolytic agent. Nucleic acid molecules of the invention are useful  
 CC for treating humans with genetically predisposed prothrombotic disorders  
 CC by gene therapy. The present sequence is human protein C derivative.  
 XX  
 SQ Sequence 419 AA:  
 Query Match 88.3%; Score 174; DB 22; Length 419;  
 Best Local Similarity 77.3%; Pred. No. 1e-20;  
 Matches 34; Conservative 0; Mismatches 10; Indels 0; Gaps 0;  
 QY 1 ANSFLXLRGSLRXRCIXICDFFXAKXIFEDVDTLAFWSKH 44  
 DB 1 ANSFLXLRGSLRXRCIEICDFFXAKXIFEDVDTLAFWSKH 44  
 RESULT 7  
 AAE08628  
 ID AAE08628 standard; Protein: 419 AA.  
 XX  
 XX AAE08628:  
 AC  
 XX  
 XX 01-NOV-2001 (first entry)  
 DT  
 XX  
 XX Human protein C derivative #2.  
 DE  
 XX Human; protein C derivative; anticoagulation activity; thrombosis;  
 KM

KM serpin inactivation; acute coronary syndrome; myocardial infarction;  
 KM vascular occlusive disorder; hypercoagulable state; angina; sepsis;  
 KM disseminated intravascular coagulation; DIC; burn; transplantation;  
 KM sickle cell disease; viral haemorrhagic fever; protein C deficiency;  
 KM haemolytic uremic syndrome; acute arterial thrombotic occlusion;  
 KM thrombocytopenia; prothrombotic disorder; gene therapy; thalassemia.  
 OS Homo sapiens.  
 XX  
 XX WO200159084-A1.  
 PN 16-AUG-2001.  
 PD  
 XX  
 XX 02-FEB-2001; 2001WO-US01221.  
 PF  
 XX 11-FEB-2000; 2000US-0181948.  
 PR 14-MAR-2000; 2000US-0189199.  
 XX  
 XX (ELIL ) LILLY & CO ELI.  
 PA  
 XX  
 XX Gerlitz BE, Grinnell BW, Jones BE;  
 PI  
 XX WPI: 2001-514662/56.  
 DR N-PSDB: AAD15226.  
 XX  
 XX Protein C derivative for treating acute coronary syndromes, vascular  
 PT occlusive disorders, thrombotic disorders and sepsis, comprises  
 PT substitutions at specified amino acid positions  
 PS  
 XX Claim 4; Page 47-48; 59pp; English.  
 XX  
 XX The invention relates to human protein C derivatives and nucleic acid  
 CC molecules encoding such derivatives. These derivatives have increased  
 CC anticoagulation activity, resistance to serpin inactivation and  
 CC increased sensitivity to thrombin activation compared to wild type  
 CC protein C, and retains the biological activity of the wild type human  
 CC protein C. Protein C derivatives are useful in the manufacture of a  
 CC medicament for the treatment of acute coronary syndromes e.g. myocardial  
 CC infarction and unstable angina; and disease states predisposing to  
 CC thrombosis; vascular occlusive disorders and hypercoagulable states e.g.  
 CC disseminated intravascular coagulation (DIC), burns, transplantations,  
 CC thalassemia, sickle cell disease, viral haemorrhagic fever and  
 CC haemolytic uremic syndrome; sepsis in combination with bacterial  
 CC permeability increasing protein; thrombotic disorders in combination  
 CC with an anti-platelet agent; protein C deficiency; acute arterial  
 CC thrombotic occlusion, thromboembolism or stenosis in coronary, cerebral  
 CC or peripheral arteries or in vascular grafts in combination with a  
 CC thrombolytic agent. Nucleic acid molecules of the invention are useful  
 CC for treating humans with genetically predisposed prothrombotic disorders  
 CC by gene therapy. The present sequence is human protein C derivative.  
 XX  
 SQ Sequence 419 AA:  
 Query Match 88.3%; Score 174; DB 22; Length 419;  
 Best Local Similarity 77.3%; Pred. No. 1e-20;  
 Matches 34; Conservative 0; Mismatches 10; Indels 0; Gaps 0;  
 QY 1 ANSFLXLRGSLRXRCIXICDFFXAKXIFEDVDTLAFWSKH 44  
 DB 1 ANSFLXLRGSLRXRCIEICDFFXAKXIFEDVDTLAFWSKH 44  
 RESULT 8  
 AAE08629  
 ID AAE08629 standard; Protein: 419 AA.  
 XX  
 XX AAE08629:  
 AC  
 XX  
 XX 01-NOV-2001 (first entry)  
 DT  
 XX  
 XX Human protein C derivative #3.  
 DE  
 XX Human; protein C derivative; anticoagulation activity; thrombosis;  
 KM

KW serpin inactivation; acute coronary syndrome; myocardial infarction;  
 KW vascular occlusive disorder; hypercoagulable state; angina; sepsis;  
 KW disseminated intravascular coagulation; DIC; burn; transplantation;  
 KW sickle cell disease; viral haemorrhagic fever; protein C deficiency;  
 KW haemolytic uraemic syndrome; acute arterial thrombotic occlusion;  
 KW thrombocytopenia; thrombotic disorder; gene therapy; thalassemia.  
 OS Homo sapiens.  
 XX  
 XX  
 FH Key Location/Qualifiers  
 FT Misc-difference 10 /note= "Encoded by CAA"  
 PN WO200159084-A1.  
 XX  
 XX 16-AUG-2001.  
 PD  
 XX  
 XX 02-FEB-2001; 2001WO-US01221.  
 PF  
 XX 11-FEB-2000; 2000US-0181948.  
 PR 14-MAR-2000; 2000US-0189199.  
 XX  
 XX (ELIL ) LULLY & CO ELI.  
 PA  
 XX Gerlitz BE, Grinnell BW, Jones BE;  
 PI WPI: 2001-514662/56.  
 XX N-PSDB: AAD15227.  
 DR  
 XX  
 XX Protein C derivative for treating acute coronary syndromes, vascular  
 PT occlusive disorders, thrombotic disorders and sepsis, comprises  
 PT substitutions at specified amino acid positions  
 XX  
 PS Claim 5; Page 48-49; 59pp; English.  
 XX  
 CC The invention relates to human protein C derivatives and nucleic acid  
 CC molecules encoding such derivatives. These derivatives have increased  
 CC anticoagulation activity, resistance to serpin inactivation and  
 CC increased sensitivity to thrombin activation compared to wild type  
 CC protein C, and retains the biological activity of the wild type human  
 CC protein C. Protein C derivatives are useful in the manufacture of a  
 CC medicament for the treatment of acute coronary syndromes e.g. myocardial  
 CC infarction and unstable angina; and disease states predisposing to  
 CC thrombosis; vascular occlusive disorders and hypercoagulable states e.g.  
 CC disseminated intravascular coagulation (DIC), burns, transplantations,  
 CC thalassemia, sickle cell disease, viral haemorrhagic fever and  
 CC haemolytic uraemic syndrome; sepsis in combination with bacterial  
 CC permeability increasing protein; thrombotic disorders in combination  
 CC with an anti-platelet agent; protein C deficiency; acute arterial  
 CC thrombotic occlusion, thromboembolism or stenosis in coronary, cerebral  
 CC or peripheral arteries or in vascular grafts in combination with a  
 CC thrombolytic agent. Nucleic acid molecules of the invention are useful  
 CC for treating humans with genetically predisposed thrombotic disorders  
 CC by gene therapy. The present sequence is human protein C derivative.  
 CC  
 XX  
 SQ Sequence 419 AA;  
 QY 1 ANSFLXXLRGSLKRCIXIXICDFXKXKXIFEDVDLAFWSKH 44  
 Db 1 ANSFLXELRHGSLERECIEICDFEAKEIFEDVDLAFWSKH 44  
 Query Match 88.3%; Score 174; DB 22; Length 419;  
 Best Local Similarity 77.3%; Pred. No. 1e-20;  
 Matches 34; Conservative 0; Mismatches 10; Indels 0; Gaps 0;

XX Human protein C derivative (S11G/032E/N33D/L194S).  
 DE  
 XX Protein C; human; coronary syndrome; thrombosis; angina;  
 KW myocardial infarction; vascular occlusive disorder;  
 KW hypercoagulability; sepsis; protein C deficiency; occlusion;  
 KW thromboembolism; stenosis; antibacterial; immunosuppressive;  
 KW thrombolytic; cardiac; antiangiogenic; anticoagulant; therapy;  
 KW mutant; mutein.  
 XX  
 XX Homo sapiens.  
 OS Synthetic.  
 XX  
 XX  
 FH Key Location/Qualifiers  
 FT Misc-difference 11  
 FT Misc-difference /note= "Ser in wild-type protein"  
 FT Misc-difference 32 /note= "Gln in wild-type protein"  
 FT Misc-difference 33 /note= "Asn in wild-type protein"  
 FT Misc-difference 194 /note= "Leu in wild-type protein"  
 FT Domain  
 FT 1..45  
 FT /note= "Gla domain"  
 FT Disulfide-bond 50..69  
 FT Disulfide-bond 59..64  
 FT Disulfide-bond 80..89  
 FT Disulfide-bond 98..109  
 FT Disulfide-bond 120..133  
 FT Disulfide-bond 141..277  
 FT Disulfide-bond 196..212  
 FT Disulfide-bond 331..345  
 FT Disulfide-bond 356..384  
 FT Disulfide-bond 156..157  
 FT /note= "cleavage makes a 2-chain inactive  
 FT precursor (155-amino acid light chain  
 FT attached via a disulfide bond to a  
 FT 262-amino acid heavy chain)"  
 FT  
 FT Modified-site 6 /note= "gamma-carboxylated"  
 FT Modified-site 7 /note= "gamma-carboxylated"  
 FT Modified-site 14 /note= "gamma-carboxylated"  
 FT Modified-site 16 /note= "gamma-carboxylated"  
 FT Modified-site 19 /note= "gamma-carboxylated"  
 FT Modified-site 20 /note= "gamma-carboxylated"  
 FT Modified-site 25 /note= "gamma-carboxylated"  
 FT Modified-site 26 /note= "gamma-carboxylated"  
 FT Modified-site 158..169 /note= "gamma-carboxylated"  
 FT Peptide  
 FT /note= "activation peptide; removal activates the  
 FT 2-chain zymogen"  
 FT Cleavage-site 169..170  
 FT /note= "thrombin cleavage site"  
 FT Modified-site 29 /note= "N-glycosylated"  
 FT Modified-site 248 /note= "N-glycosylated"  
 FT Modified-site 313 /note= "N-glycosylated"  
 FT Modified-site 329 /note= "N-glycosylated"  
 FT /note= "N-glycosylated"  
 XX  
 XX WO200157193-A2.  
 XX  
 XX 09-AUG-2001.  
 PD  
 XX  
 XX 19-JAN-2001; 2001WO-US000020.

XX 02-FEB-2000; 2000US-0179801.  
 PR 14-MAR-2000; 2000US-0189197.  
 XX (ELIL ) LILLY & CO ELI.  
 PA Gerlitz BE, Jones BE;  
 PI  
 XX  
 DR WPI: 2001-496919/54.  
 N-PSDB: AAH26363.  
 XX  
 PT Novel human protein C derivative for treating, e.g., myocardial  
 PT infarction, unstable angina, sepsis, thrombotic disorders, acute  
 PT arterial thrombotic occlusion, and thromboembolism -  
 XX  
 PS Claim 3; Page 52-53; 63pp; English.  
 XX  
 CC The present sequence is that of a claimed human protein C  
 CC derivative in which Ser at amino acid position 11 of the mature  
 CC wild-type protein C sequence (see AAB82673) is substituted with  
 CC Gly, Gln at position 32 with Glu, Asn at position 33 with Asp, and  
 CC Leu at position 194 with Ser. The protein is an example of protein  
 CC derivatives of the invention that have at least 2 amino acid  
 CC substitutions, but which have increased anticoagulant activity and  
 CC resistance to inactivation by serpins compared with the wild-type  
 CC protein. A method of producing the biological activity of the wild-type  
 CC protein. A method of producing the derivatives using recombinant  
 CC DNA methods is claimed. The protein C derivatives are useful for  
 CC treating coronary syndromes and disease states predisposing to  
 CC thrombosis (e.g., myocardial infarction and unstable angina),  
 CC vascular occlusive disorders and hypercoagulable states, sepsis (in  
 CC combination with bactericidal permeability increasing protein or  
 CC with tissue factor pathway inhibitor), thrombotic disorders (in  
 CC combination with an anti-platelet agent or by local delivery through  
 CC an intracoronary catheter), protein C deficiency, acute arterial  
 CC thrombotic occlusion, thromboembolism, or stenosis in coronary,  
 CC cerebral or peripheral arteries or in vascular grafts. Human  
 CC patients with genetically predisposed prothrombotic disorders may  
 CC be treated by gene therapy (all claimed).  
 XX  
 SQ Sequence 419 AA:  
 Query Match 88.3%; Score 174; DB 22; Length 419;  
 Best Local Similarity 77.3%; Pred. No. 1e-20;  
 Matches 34; Conservative 0; Mismatches 10; Indels 0; Gaps 0;  
 QY 1 ANSFLLXLRGSLXKXCIXICPFXXAKXIFEDVDOTLAWSKH 44  
 1 ANSFLELRHGSLEKCEIEICDFEAKKEIFEDVDTLAWSKH 44  
 DB  
 RESULT 10  
 AAB82676  
 ID AAB82676 standard; Protein; 419 AA.  
 XX  
 AC AAB82676;  
 XX  
 DT 15-OCT-2001 (first entry)  
 XX  
 DE Human protein C derivative (S11G/Q32E/N33D/L194S/T254S).  
 XX  
 KW Protein C; human; coronary syndrome; thrombosis; angina;  
 KW myocardial infarction; vascular occlusive disorder;  
 KW hypercoagulation; sepsis; protein C deficiency; occlusion;  
 KW thromboembolism; stenosis; antibacterial; immunosuppressive;  
 KW thrombolytic; cardiant; antilanginal; anticoagulant; therapy;  
 KW mutant; mutain.  
 XX  
 OS Homo sapiens.  
 OS Synthetic.  
 XX  
 FH Key Location/Qualifiers  
 FT Misc-difference 11

FT /note- "Ser in wild-type protein"  
 FT 32 /note- "Gln in wild-type protein"  
 FT Misc-difference 33 /note- "Asn in wild-type protein"  
 FT 194 /note- "Leu in wild-type protein"  
 FT Misc-difference 254 /note- "Thr in wild-type protein"  
 FT 1..45 /note- "Gla domain"  
 FT 50..69 /note- "Gla domain"  
 FT 59..64 Disulfide-bond  
 FT 80..89 Disulfide-bond  
 FT 98..109 Disulfide-bond  
 FT 120..133 Disulfide-bond  
 FT 141..277 Disulfide-bond  
 FT 196..212 Disulfide-bond  
 FT 331..345 Disulfide-bond  
 FT 356..384 Disulfide-bond  
 FT 156..157 Cleavage-site  
 FT /note- "cleavage makes a 2-chain inactive  
 precursor (135-amino acid light chain  
 attached via a disulfide bond to a  
 262-amino acid heavy chain)"  
 FT Modified-site 6 /note- "gamma-carboxylated"  
 FT 7 /note- "gamma-carboxylated"  
 FT Modified-site 14 /note- "gamma-carboxylated"  
 FT Modified-site 16 /note- "gamma-carboxylated"  
 FT Modified-site 19 /note- "gamma-carboxylated"  
 FT Modified-site 20 /note- "gamma-carboxylated"  
 FT Modified-site 25 /note- "gamma-carboxylated"  
 FT Modified-site 26 /note- "gamma-carboxylated"  
 FT Modified-site 158..169 /note- "gamma-carboxylated"  
 FT Peptide /note- "activation peptide; removal activates the  
 2-chain zymogen"  
 FT Cleavage-site 169..170 /note- "thrombin cleavage site"  
 FT Modified-site 29 /note- "N-glycosylated"  
 FT Modified-site 248 /note- "N-glycosylated"  
 FT Modified-site 313 /note- "N-glycosylated"  
 FT Modified-site 329 /note- "N-glycosylated"  
 FT Modified-site /note- "N-glycosylated"  
 XX  
 PN MO200157193-A2.  
 XX  
 PD 09-AUG-2001.  
 XX  
 PE 19-JAN-2001; 2001MO-US00020.  
 XX  
 XX 02-FEB-2000; 2000US-0179801.  
 PR 14-MAR-2000; 2000US-0189197.  
 XX (ELIL ) LILLY & CO ELI.  
 PA Gerlitz BE, Jones BE;  
 PI  
 XX  
 DR WPI: 2001-496919/54.  
 DR N-PSDB: AAH26364.  
 XX  
 PT Novel human protein C derivative for treating, e.g., myocardial  
 PT infarction, unstable angina, sepsis, thrombotic disorders, acute

PT arterial thrombotic occlusion, and thromboembolism -

PS Claim 4; Page 53-54; 63pp; English.

XX

XX The present sequence is that of a claimed human protein C derivative

CC in which Ser at position 11 of the mature wild-type protein C

CC sequence (see AbB82673) is substituted with Gly, Gln at position 32

CC with Glu, Asn at position 33 with Asp, Leu at position 194 with Ser,

CC and Thr at position 254 with Ser. It is an example of protein C

CC derivatives of the invention that have at least 2 amino acid

CC substitutions, but which have increased anticoagulant activity and

CC resistance to inactivation by serpins compared with the wild-type

CC protein, while retaining the biological activity of the wild-type

CC protein. A method of producing the derivatives using recombinant

CC DNA methods is claimed. The protein C derivatives are useful for

CC treating coronary syndromes and disease states predisposing to

CC thrombosis (e.g. myocardial infarction and unstable angina),

CC vascular occlusive disorders and hypercoagulable states, sepsis (in

CC combination with bactericidal permeability increasing protein or

CC with tissue factor pathway inhibitor), thrombotic disorders (in

CC combination with an anti-platelet agent or by local delivery through

CC an intracoronary catheter), protein C deficiency, acute arterial

CC thrombotic occlusion, thromboembolism, or stenosis in coronary,

CC cerebral or peripheral arteries or in vascular grafts. Human

CC patients with genetically predisposed prothrombotic disorders may

CC be treated by gene therapy (all claimed).

XX

SO Sequence 419 AA:

Query Match 88.3%; Score 174; DB 22; Length 419;

Best Local Similarity 77.3%; Pred. No. 1e-20;

Matches 34; Conservative 0; Mismatches 10; Indels 0; Gaps 0;

OY 1 ANSFLXLRQGS�XRXCIXICDFFXAKXIFEDVDDTLAFMSKH 44

||||| ||||| ||||| ||||| ||||| ||||| ||||| ||||| |||||

DB 1 ANSFLFLRHGSLRECEIEICDFEAKEIFEDVDDTLAFMSKH 44

RESULT 11

AAV18298 ID AAV18298 standard; peptide; 44 AA.

XX

XX AAV18298;

AC

XX

DT 17-AUG-1999 (first entry)

XX

DE Modified GLA domain of vitamin K-dependent protein.

XX

KW GLA domain; muten; vitamin K-dependent protein; clotting disorder;

KM therapy.

XX

XX Homo sapiens.

OS Synthetic.

OS

XX

XX Key Location/Qualifiers

FT Misc-difference 1..44

FT /note= "Xaa= gamma-carboxyglutamic acid, or glutamic

FT acid"

FT

XX

XX W09920767-A1.

PN

XX

XX 29-APR-1999.

PD

XX

XX 20-OCT-1998; 98WO-US22152.

PF

XX

PR 23-OCT-1997; 97US-0955636.

XX

XX (MINU ) UNIV MINNESOTA.

PA

XX

PI Neisestuen GL;

XX

XX WPI; 1999-288309/24.

DR

XX

PT Vitamin K-dependent polypeptide with modified gamma-carboxyglutamic

PT acid domain, useful for treating clotting disorders

XX

XX

PS Claim 7; Page 78; 86pp; English.

XX

XX This sequence represents a modified GLA (gamma-carboxyglutamic acid)

CC domain. The invention relates to a vitamin K-dependent polypeptide

CC comprising a modified GLA domain containing an amino acid substitution

CC which enhances membrane binding of the modified polypeptide as compared

CC to the native polypeptide. The polypeptide is used to treat a clotting

CC disorder by decreasing or increasing clot formation. Modification of the

CC GLA domain results in a protein which has enhanced membrane binding

CC affinity as compared to the native protein.

XX

SO Sequence 44 AA:

Query Match 87.8%; Score 173; DB 20; Length 44;

Best Local Similarity 97.7%; Pred. No. 1.3e-21;

Matches 43; Conservative 0; Mismatches 1; Indels 0; Gaps 0;

OY 1 ANSFLXLRQGS�XRXCIXICDFFXAKXIFEDVDDTLAFMSKH 44

||||| ||||| ||||| ||||| ||||| ||||| ||||| ||||| |||||

DB 1 ANSFLXLRQGS�XRXCIXICDFFXAKXIFEDVDDTLAFMSKH 44

RESULT 12

AAV18299 ID AAV18299 standard; peptide; 44 AA.

XX

XX AAV18299;

AC

XX

DT 17-AUG-1999 (first entry)

XX

DE Modified GLA domain of vitamin K-dependent protein.

XX

KW GLA domain; muten; vitamin K-dependent protein; clotting disorder;

KM therapy.

XX

XX Homo sapiens.

OS Synthetic.

OS

XX

XX Key Location/Qualifiers

FT Misc-difference 1..44

FT /note= "Xaa= gamma-carboxyglutamic acid, or glutamic

FT acid"

FT

XX

XX W09920767-A1.

PN

XX

XX 29-APR-1999.

PD

XX

XX 20-OCT-1998; 98WO-US22152.

PF

XX

PR 23-OCT-1997; 97US-0955636.

XX

XX (MINU ) UNIV MINNESOTA.

PA

XX

PI Neisestuen GL;

XX

XX WPI; 1999-288309/24.

DR

XX

XX Claim 8; Page 78; 86pp; English.

XX

XX This sequence represents a modified GLA (gamma-carboxyglutamic acid)

CC domain. The invention relates to a vitamin K-dependent polypeptide

CC comprising a modified GLA domain containing an amino acid substitution

CC which enhances membrane binding of the modified polypeptide as compared

CC to the native polypeptide. The polypeptide is used to treat a clotting

CC disorder by decreasing or increasing clot formation. Modification of the

CC GLA domain results in a protein which has enhanced membrane binding

CC affinity as compared to the native protein.





OS Synthetic.  
 XX  
 FH Key Location/Qualifiers  
 FT Misc-difference 1..44  
 FT /note- "Xaa- gamma-carboxyglutamic acid, or glutamic  
 FT acid"  
 XX  
 XX W09920767-A1.  
 PN  
 XX 29-APR-1999.  
 PD  
 XX 20-OCT-1998; 98WO-US22152.  
 PF  
 XX 23-OCT-1997; 97US-0955636.  
 PR  
 XX (MINU ) UNIV MINNESOTA.  
 PA  
 XX Nelsetuen GL;  
 PI  
 XX WPI; 1999-288309/24.  
 DR  
 XX  
 XX Vitamin K-dependent polypeptide with modified gamma-carboxyglutamic  
 PT acid domain, useful for treating clotting disorders  
 XX  
 PS Disclosure: Page 79-80; 86pp; English.  
 XX  
 CC This sequence represents a modified GLA (gamma-carboxyglutamic acid)  
 CC domain. The invention relates to a vitamin K-dependent polypeptide  
 CC comprising a modified GLA domain containing an amino acid substitution  
 CC which enhances membrane binding of the modified polypeptide as compared  
 CC to the native polypeptide. The polypeptide is used to treat a clotting  
 CC disorder by decreasing or increasing clot formation. Modification of the  
 CC GLA domain results in a protein which has enhanced membrane binding  
 CC affinity as compared to the native protein.  
 XX  
 SQ Sequence 44 AA;

Query Match 81.2%; Score 160; DB 20; Length 44;  
 Best Local Similarity 93.2%; Pred. No. 2e-19;  
 Matches 41; Conservative 0; Mismatches 3; Indels 0; Gaps 0;

OY 1 ANSFLXXLRHSSSLKXCIXXICDFXXAKXIFEDVDDTLAFWSKH 44  
 ||||||| ||||||||||||||| ||||||||||||  
 Db 1 ANSFLXXLRHSSSLKXCIXXICDFXXAKXIFEDVDDTLAFWSKH 44

Search completed: May 16, 2003, 10:14:32  
 Job time : 36 secs

GenCore version 5.1.4.p5.4578  
Copyright (c) 1993 - 2003 CompuGen Ltd.

OM protein - protein search, using sw model

Run on: May 16, 2003, 10:12:44 : Search time 18 seconds

(without alignments)  
234.995 Million cell updates/sec

Title: SEQ1-4EDITS

Perfect score: 197  
Sequence: 1 ANSFLXHLRGLSLXRCIXX.....XXAKXIFedVDTLAFMSKH 44

Scoring table: BLOSUM62  
Gapop 10.0, Capext 0.5

Searched: 283224 seqs, 96134422 residues

Total number of hits satisfying chosen parameters: 283224

Minimum DB seq length: 0  
Maximum DB seq length: 200000000

Post-processing: Minimum Match 0%

Listing first 45 summaries

Database :

1: PIR\_73:.\*  
2: PIR1:.\*  
3: PIR2:.\*  
4: PIR3:.\*

Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

#### SUMMARIES

Result No.	Score	Query Match	Length	ID	Description
1	160	81.2	461	1 KXHU	protein C (activat
2	140	71.1	461	1 JX0210	protein C (activat
3	139	70.6	461	1 S18994	protein C (activat
4	122	61.9	456	1 KXBO	protein C (activat
5	115	58.4	482	1 EXRT	coagulation factor
6	114	57.9	492	1 EXBO	coagulation factor
7	110	55.8	488	1 EXHU	coagulation factor
8	101	51.3	443	2 I46932	coagulation factor
9	99	50.3	466	1 KFH07	coagulation factor
10	86.5	43.9	617	2 S10511	thrombin (EC 3.4.2
11	86.5	43.9	618	2 A35827	thrombin (EC 3.4.2
12	86	43.7	475	1 EXCH	coagulation factor
13	85	43.1	407	1 KEB07	coagulation factor
14	85	43.1	642	2 S53434	plasma protein S p
15	85	43.1	676	1 KXHU5	plasma protein S p
16	84	42.6	622	1 T8HU	thrombin (EC 3.4.2
17	84	41.1	646	2 S38819	thrombin (EC 3.4.2
18	80	40.6	452	1 A30351	coagulation factor
19	80	40.6	459	2 JQ0419	coagulation factor
20	80	40.6	461	1 KFH0	coagulation factor
21	80	40.6	675	1 KXBO5	plasma protein S p
22	78	39.6	642	2 S53433	plasma protein S p
23	78	39.6	675	1 KXRT5	plasma protein S p
24	73	37.1	416	1 KFB0	coagulation factor
25	72	36.5	625	1 T8BO	thrombin (EC 3.4.2
26	71	36.0	675	1 KXMS	plasma protein S p
27	69.5	35.3	396	1 KXBO2	plasma protein S p
28	65.5	33.2	422	1 KXHU2	plasma protein Z p
29	65	33.0	673	2 A48089	growth arrest-spec

30	64	32.5	674	2 I55476	growth potentialin
31	63	32.0	678	2 B48089	growth arrest-spec
32	56.5	28.7	594	2 D84859	probable MAP kinase
33	54.5	27.7	603	2 C96575	probable MAP kinase
34	53.5	27.2	576	2 G96763	probable MAP kinase
35	53	26.9	606	2 T40556	hypothetical prote
36	50	25.4	1684	2 T02367	hypothetical prote
37	49	24.9	1363	2 T58375	protein-tyrosine k
38	48.5	24.6	323	2 T25948	hypothetical prote
39	48.5	24.6	510	2 E82918	ammonium transport
40	48	24.4	422	2 T39306	mitogen-activated
41	48	24.4	1235	2 D32433	VSG expression sit
42	48	24.4	1298	2 A48999	protein-tyrosine k
43	47.5	24.1	1089	1 S33727	platelet-derived g
44	47	23.9	182	2 JCI189	tyrosine kinase re
45	47	23.9	245	1 NDECR5	type II site-speci

#### ALIGNMENTS

RESULT 1  
KXHU  
protein C (activated) (EC 3.4.21.69) precursor - human  
N:Alternate names: autoprothrombin IIA; plasma protein C  
C:Species: Homo sapiens (man)  
C>Date: 17-Mar-1987 #sequence, revision 17-Mar-1987 #text, change 16-Jul-1999  
C:Accession: A22331; A25426; A21781; A23789; A00927  
R:Foster, D.C.; Yoshitake, S.; Davie, E.W.  
Proc. Natl. Acad. Sci. U.S.A. 82, 4673-4677, 1985  
A:Title: The nucleotide sequence of the gene for human protein C.  
A:Reference number: A22331; MUID:85270390; PMID:2991887  
A:Accession: A22331  
A:Molecule type: DNA  
A:Residues: 1-461 <F0S1>  
A:Cross-references: GB:M11228; NID:g190333; PIDN:AAA60166.1; PID:g190334  
R:Plutsky, J.; Hoskins, J.A.; Long, G.L.; Crabtree, G.R.  
Proc. Natl. Acad. Sci. U.S.A. 83, 546-550, 1986  
A:Title: Evolution and organization of the human protein C gene.  
A:Reference number: A25426; MUID:86120978; PMID:3511471  
A:Accession: A25426  
A:Molecule type: DNA  
A:Residues: 1-445, 1-446-461 <PLU>  
A:Cross-references: GB:M12712; NID:g190330; PIDN:AAA60165.1; PID:g190332  
R:Foster, D.; Davie, E.W.  
Proc. Natl. Acad. Sci. U.S.A. 81, 4766-4770, 1984  
A:Title: Characterization of a cDNA coding for human protein C.  
A:Reference number: A21781; MUID:84272714; PMID:6589623  
A:Accession: A21781  
A:Molecule type: mRNA  
A:Residues: Q', 107-461 <F0S2>  
A:Cross-references: GB:X02750; NID:g35689; PIDN:CAA26528.1; PID:g763120  
R:Meleisch, J.P.; Broze Jr., G.J.  
J. Biol. Chem. 265, 11397-11404, 1990  
A:Title: Beta protein C is not glycosylated at asparagine 329. The rate of translati  
A:Reference number: A44605; MUID:90239094; PMID:1694179  
A:Contents: annotation: carbohydrate binding sites; activation peptide  
A:Note: the alpha form of protein C is glycosylated at Asn-329, and the beta form is  
J. Biol. Chem. 267, 5102-5107, 1992  
A:Title: O-linked fucose is present in the first epidermal growth factor domain of f  
A:Reference number: A44606; MUID:92184750; PMID:1544884  
A:Contents: annotation: beta-hydroxyspartic acid  
A:Comment: protein C is the zymogen of the vitamin K-dependent serine proteinase tha  
ivation of factor Va is strongly enhanced by complexing with protein S. Protein C al

C:Comment: Protein C is synthesized in the liver as a single chain precursor, which is then cleaved into two chains, which cleaves a dodecapeptide from the amino end of the heavy chain; this reaction, C:Genetics:

A:Gene: GDB:PROC

A:Cross-references: GDB:120317; OMIM:176860

A:Map position: 2q13-2q21

A:Introns: 24/1: 79/3: 88/1: 134/1: 179/1: 226/3: 266/1:

C:Superfamily: coagulation factor X; EGF homology; Gla domain homology; trypsin homology

C:Keywords: anticoagulant; beta-hydroxyaspartic acid; blood coagulation; calcium binding

F:1-32/Domain: signal sequence #status predicted <SIG>

F:27-86/Domain: Gla domain homology <Gla>

F:33-42/Domain: propeptide #status predicted <PRO>

F:43-197/Product: protein C light chain #status predicted <LCH>

F:92-131/Domain: EGF homology <EG1>

F:140-115/Domain: EGF homology <EG2>

F:200-461/Product: protein C heavy chain #status predicted <HCH>

F:200-211/Domain: activation peptide #status experimental <AP>

F:212-445/Domain: trypsin homology <TRY>

F:48-49, 56, 58, 61, 62, 67, 68, 71/Modified site: gamma-carboxyglutamic acid (Glu) #status exp

F:59-64, 92-105, 101-120, 122-131, 140-151, 147-160, 162-175, 183-319, 238-254, 373-387, 398-426/T

F:106-111/Disulfide bonds: #status predicted

F:110/Binding site: carbonylate (Thr) (covalent) #status absent

F:113/Modified site: erythro-beta-hydroxyaspartic acid (Asp) #status experimental

F:139, 290, 355/Binding site: carbonylate (Asn) (covalent) #status experimental

F:211-212/Cleavage site: Arg-Ileu (thrombin) #status experimental

F:253, 299, 402/Active site: His, Asp, Ser #status predicted

F:371/Binding site: carbonylate (Asn) (covalent) (partial) #status atypical

Query Match 81.2%; Score 160; DB 1; Length 461;

Best Local Similarity 70.5%; Pred. No. 8, 8e-18;

Matches 31; Conservative 2; Mismatches 11; Indels 0; Gaps 0;

Oy 1 ANSFLXLRGSLKRCIXICDFYXAKXIFEDVDTLAFNSKH 44

Db 43 ANSFLERHSLRECEIEICDFEAEKIFQNVDDTLAFNSKH 86

RESULT 2

JX0210

Protein C (activated) (EC 3.4.21.69) precursor - mouse

N:Alternate names: vitamin K-dependent serine proteinase

C:Species: Mus musculus (house mouse)

C:Date: 10-Sep-1999 #sequence\_revision 10-Sep-1999 #text\_change 16-Jun-2000

R:Trada, N.; Sato, M.; Tsujimura, A.; Iwase, R.; Hashimoto-Gotoh, T.

J. Biochem. 111, 491-495, 1992

A:Title: Isolation and characterization of a mouse protein C cDNA.

A:Reference number: JX0210; MUID:92316897; PMID:1618739

A:Accession: JX0210

A:Molecule type: mRNA

A:Residues: 1-461 <TAD>

A:Cross-references: GB:DL0445; NID:9220385; PIDN:BA01235.1; PID:9220386

A:Experimental source: liver

C:Comment: Protein C is the zymogen of the vitamin K-dependent serine proteinase that re

S:

C:Superfamily: coagulation factor X; EGF homology; Gla domain homology; trypsin homology

C:Keywords: beta-hydroxyaspartic acid; blood coagulation; calcium binding; carboxylut

F:1-33/Domain: signal sequence #status predicted <SIG>

F:27-85/Domain: Gla domain homology <Gla>

F:34-41/Domain: propeptide #status predicted <PRO>

F:42-196, 199-461/Product: protein C #status predicted <PRC>

F:42-196/Domain: light chain #status predicted <PCL>

F:91-130/Domain: EGF homology <EG1>

F:139-174/Domain: EGF homology <EG2>

F:199-461/Domain: heavy chain #status predicted <PCH>

F:199-211/Domain: activation peptide #status predicted <ACT>

F:212-461/Product: vitamin K-dependent serine proteinase #status predicted <VIT>

F:212-445/Domain: trypsin homology <TRY>

F:47, 48, 55, 57, 60, 61, 66, 67, 70, 76/Modified site: gamma-carboxyglutamic acid (Glu) #status

F:112/Modified site: erythro-beta-hydroxyaspartic acid (Asp) #status predicted

F:121-130, 139-150, 146-159, 161-174, 182-319, 238-254, 373-387, 398-426/Disulfide bonds: #stat

F:214, 290, 355/Binding site: carbonylate (Asn) (covalent) #status predicted

F:253, 299, 402/Active site: His, Asp, Ser #status predicted

Query Match 71.1%; Score 140; DB 1; Length 461;

Best Local Similarity 59.1%; Pred. No. 1, 5e-14;

Matches 26; Conservative 7; Mismatches 11; Indels 0; Gaps 0;

Oy 1 ANSFLXLRGSLKRCIXICDFYXAKXIFEDVDTLAFNSKH 44

Db 42 ANSFLERHSLRECEIEICDFEAEKIFQNVDDTLAFNSKH 85

RESULT 3

S18994

protein C (activated) (EC 3.4.21.69) precursor - rat

C:Species: Rattus norvegicus (Norway rat)

C:Date: 10-Sep-1999 #sequence\_revision 10-Sep-1999 #text\_change 29-Oct-1999

C:Accession: S18994; S24312

R:Okafuji, T.; Maekawa, K.; Nawa, K.; Marumoto, Y.

submitted to the EMBL Data Library, February 1992

A:Description: The cDNA cloning and mRNA expression of rat protein C.

A:Reference number: S18994

A:Accession: S18994

A:Status: preliminary

A:Molecule type: mRNA

A:Residues: 1-461 <OKA>

A:Cross-references: EMBL:X64336; NID:956962; PIDN:CAA45617.1; PID:956963

R:Okafuji, T.; Maekawa, K.; Nawa, K.; Marumoto, Y.

Biochim. Biophys. Acta 1131, 329-332, 1992

A:Title: The cDNA cloning and mRNA expression of rat protein C.

A:Reference number: S24312; MUID:92329550; PMID:1627650

A:Accession: S24312

A:Status: preliminary

A:Molecule type: mRNA

A:Residues: 1-461 <OKA2>

A:Cross-references: EMBL:X64336; NID:956962; PIDN:CAA45617.1; PID:956963

C:Superfamily: coagulation factor X; EGF homology; Gla domain homology; trypsin homol

C:Keywords: beta-hydroxyaspartic acid; glycoprotein; hydrolase; serine proteinase

F:1-32/Domain: signal sequence #status predicted <SIG>

F:27-85/Domain: Gla domain homology <Gla>

F:33-42/Domain: propeptide #status predicted <PRO>

F:43-461/Product: protein C #status predicted <PRC>

F:91-130/Domain: EGF homology <EG1>

F:139-174/Domain: EGF homology <EG2>

F:213-445/Domain: trypsin homology <TRY>

F:47, 48, 55, 57, 60, 61, 66, 67, 70, 76/Modified site: gamma-carboxyglutamic acid (Glu) #stat

F:112/Modified site: erythro-beta-hydroxyaspartic acid (Asp) #status predicted

F:121-130, 139-150, 146-159, 161-174, 182-320, 239-255, 373-387, 398-426/Disulfide bonds: #s

F:215, 291, 355/Binding site: carbonylate (Asn) (covalent) #status predicted

F:254, 300, 402/Active site: His, Asp, Ser #status predicted

Query Match 70.6%; Score 139; DB 1; Length 461;

Best Local Similarity 59.1%; Pred. No. 2, 1e-14;

Matches 26; Conservative 7; Mismatches 11; Indels 0; Gaps 0;

Oy 1 ANSFLXLRGSLKRCIXICDFYXAKXIFEDVDTLAFNSKH 44

Db 42 ANSFLERHSLRECEIEICDFEAEKIFQNVDDTLAFNSKH 85

RESULT 4

KXBO

Protein C (activated) (EC 3.4.21.69) precursor - bovine (fragment)

N:Alternate names: autoproteolysin IIA; plasma protein C

C:Species: Bos primigenius taurus (cattle)

C:Date: 30-Nov-1980 #sequence\_revision 17-Mar-1987 #text\_change 16-Jul-1999

C:Accession: A26250; A18385; A18386; A00928

R:Long, G.L.; Balagaje, R.M.; MacGillivray, R.T.A.

Proc. Natl. Acad. Sci. U.S.A. 81, 5653-5656, 1984

A:Title: Cloning and sequence of liver cDNA coding for bovine protein C.

A:Reference number: A26250; MUID:85014826; PMID:6091100

A:Accession: A26250

A:Molecule type: mRNA

A:Residues: 1-456 <ION>

R:Fernlund, P.; Stenflo, J.

J. Biol. Chem. 257, 12170-12179, 1982  
 A:Title: Amino acid sequence of the light chain of bovine protein C.  
 A:Reference number: A18385; MUID:83007325; PMID:6896876  
 A:Accession: A18385  
 A:Molecule type: protein  
 A:Residues: 40-194 <PER>  
 A>Note: 82-Lys was also found  
 R:Drakenberg, T.; Fernlund, P.; Koepstorff, P.; Stenflo, J.  
 Proc. Natl. Acad. Sci. U.S.A. 80, 1802-1806, 1983  
 A:Title: beta-Hydroxyaspartic acid in vitamin K-dependent protein C.  
 A:Reference number: A19316; MUID:83169769; PMID:6572939  
 A:Contents: annotation: revision to residue 110  
 R:Stenflo, J.; Fernlund, P.  
 J. Biol. Chem. 257, 12180-12190, 1982  
 A:Title: Amino acid sequence of the heavy chain of bovine protein C.  
 A:Reference number: A18386; MUID:83007326; PMID:6896877  
 A:Accession: A18386  
 A:Molecule type: protein  
 A:Residues: 197-454, 'pv' <SHE>  
 R:Esmon, N.L.; Debaule, L.E.; Esmon, C.T.  
 J. Biol. Chem. 258, 5548-5553, 1983  
 A:Title: Proteolytic formation and properties of gamma-carboxyglutamic acid-domainless protein C.  
 A:Reference number: A37541; MUID:83213513; PMID:6304092  
 A:Contents: annotation: activation; calcium binding  
 R:Johnson, A.E.; Esmon, N.L.; Lane, T.M.; Esmon, C.T.  
 J. Biol. Chem. 258, 5554-5560, 1983  
 A:Title: Structural changes required for activation of protein C are induced by Ca2+ binding.  
 A:Reference number: A37542; MUID:83213514; PMID:6406503  
 A:Contents: annotation: activation; calcium binding  
 C:Comment: protein C is the zymogen of the vitamin K-dependent serine protease that releases active protein C.  
 S:Comment: protein C is synthesized in the liver as a single chain precursor, which is a dimer, which cleaves a tetradecapeptide from the amino end of the heavy chain; this reaction is catalyzed by the gamma-carboxyglutamic acid (Gla) residues and, with strict cognation of the thrombin-thrombomodulin complex.  
 C:Comment: The gamma-carboxyglutamic acid residues arise by a posttranslational, vitamin K-dependent modification of the glutamic acid residues.  
 C:Superfamily: coagulation factor X; EGF homology; Gla domain homology; trypsin homology  
 C:Keywords: anticoagulant; beta-hydroxyaspartic acid; blood coagulation; calcium binding  
 F:1-29/Domain: signal sequence (fragment) #status predicted <SIG>  
 F:24-83/Domain: Gla domain homology <GLA>  
 F:30-39/Domain: propeptide #status predicted <PRO>  
 F:40-194/Product: protein C light chain #status experimental <LCH>  
 F:98-128/Domain: EGF homology <EGF>  
 F:137-172/Domain: EGF homology <EGF>  
 F:197-456/Product: protein C heavy chain #status experimental <HCH>  
 F:197-210/Domain: activation peptide #status experimental <AP>  
 F:211-440/Domain: trypsin homology <TRY>  
 F:45-46, 53-55, 58-59, 62, 64, 65, 68, 74/Modified site: gamma-carboxyglutamic acid (Glu) #status predicted  
 F:110/Modified site: gamma-carboxyglutamic acid (Glu) #status predicted  
 F:119-128, 137-148, 144-157, 159-172, 180-318, 237-253, 368-382, 393-421/Disulfide bonds: #status predicted  
 F:136, 289, 350/Binding site: carboxyhydrate (Asn) (covalent) #status predicted  
 F:257, 298, 397/Active site: His, Asp, Ser #status predicted  
 F:366/Binding site: carboxyhydrate (Asn) (covalent) #status predicted

A:Accession: S49075  
A:Molecule type: mRNA  
A:Residues: 1-482 <STAL>  
A:Cross-references: EMBL:X79807; NID:g506600; PIDN:CAA56202.1; PID:g506601  
A>Note: submitted to the EMBL Data Library, June 1994  
R>Note: neither the complete nucleic acid sequence nor the complete translation are s  
R:Stanton, C.; Ross, R.P.; Hutson, S.; Wallin, R.  
Gene 169, 269-273, 1996  
A:Title: Processing and expression of rat and human clotting factor-X-encoding cDNAs.  
A:Reference number: J04670; MUID:96194815; PMID:8647460  
A:Accession: J04670  
A:Molecule type: mRNA  
A:Residues: 1-482 <STAZ>  
A:Cross-references: EMBL:X79807; NID:g506600; PIDN:CAA56202.1; PID:g506601  
A:Experimental source: Cos-1 cell  
R:Enyoji, K.; Miyazaki, K.; Kato, H.  
J. Biochem. 109, 890-898, 1991  
A:Title: Characterization of rat factors X and Xa: demonstration of factor Xa in rat  
A:Reference number: PS0190; MUID:92041742; PMID:11718949  
A:Accession: PS0191  
A:Molecule type: protein  
A:Residues: 41-58, 'X', 60-65 <ENJ1>  
A:Accession: PS0190  
A:Molecule type: protein  
A:Residues: 183-186, 'X', 188-207 <ENU2>  
R:Murakawa, M.; Okamura, T.; Kamura, T.; Kuroiwa, M.; Harada, M.; Niho, Y.  
Eur. J. Haematol. 52, 162-168, 1994  
A:Title: Analysis of the partial nucleotide sequences and deduced primary structures  
A:Reference number: I46196; MUID:94222160; PMID:8168596  
A:Accession: 162745  
A>Status: preliminary; translated from GB/EMBL/DDBJ  
A:Molecule type: DNA  
A:Residues: 295-383, 'G', 385-455 <MUR>  
A:Cross-references: GB:D21215; NID:g415309; PIRN:BAA04756.1; PID:g455396  
C:Function:  
A:Description: catalyzes the proteolytic activation of prothrombin to thrombin in the  
A:Pathway: blood coagulation  
C:Superfamily: coagulation factor X; EGF homology; Gla domain homology; trypsin homo]  
C:Keywords: beta-hydroxyaspartic acid; blood coagulation; calcium binding; carboxylic  
F:1-23/Domain: signal sequence #status predicted <SIG>  
F:24-40/Domain: propeptide #status predicted <PRO>  
E:25-84/Domain: Gla domain homology <GLA>  
F:81-179/Product: coagulation factor X light chain #status predicted <LCB>  
F:90-121/Domain: EGF homology <EGF>  
F:129-164/Domain: EGF homology <EG2>  
F:183-482/Product: coagulation factor X heavy chain #status predicted <HCH>  
F:183-231/Domain: activation peptide #status predicted <APT>  
F:232-482/Product: coagulation factor Xa heavy chain #status predicted <ACT>  
E:233-460/Domain: trypsin homology <TRY>  
F:446-47, 54, 56, 59, 60, 65, 69, 72, 79/Modified site: gamma-carboxyglutamic acid (Glu) #  
E:57-62, 90-101, 95-110, 112-121, 129-140, 136-149, 151-164, 172-340, 238-243, 259-275, 388-402  
F:103/Modified site: erythro-beta-hydroxyaspartic acid (Asp) #status predicted  
F:187/Binding site: carboxylate (Asn) (covalent) #status experimental  
E:208/Binding site: carboxylate (Thr) (covalent) #status predicted  
F:218/Binding site: carboxylate (Asn) (covalent) #status predicted  
F:231-232/Cleavage site: Arg-Ile (coagulation factor IXa, coagulation factor VIIa) #  
F:274, 320, 417/Active site: His, Asp, Ser #status predicted

Query Match 58.4% Score 115; DB 1; Length 482;  
Best Local Similarity 43.2%; Pred. NO. 1.7e-10;  
Matches 19; Conservative 10; Mismatches 15; Indels 0; Gaps 0;

OY 1 ANSEFLXLROSLKRCXCIXICDFXXAKXIREFDDVLAFMSKH 44  
|||||::||| | : ||| : ||| :  
Db 41 ANSFEELIKKNLEKRECEVEELCSFEFAREVDEDEKTEPFANKY 84

RESULT 6  
EXBO  
coagulation factor Xa (EC 3.4.21.6) precursor - bovine  
N:Alternate names: Stuart factor  
C:Species: Bos primigenius taurus (cattle)  
[Date: 24-Apr-1994 #sequence\_revision 17-Mar-1987 #text\_change 16-Jul-1999]

C:Accession: A22867; A14997; A12030; A34412; S39414; A00925  
 R:Funf, M.R.; Campbell, R.M.; MacGillivray, T.A.  
 Nucleic Acids Res. 12, 4461-4492, 1984  
 A:Title: Blood coagulation factor X mRNA encodes a single polypeptide chain containing a  
 A:Reference number: A22867; MUID:84247315; PMID:6330671  
 A:Accession: A22867  
 A:Molecule type: mRNA  
 A:Residues: 1-487 <FUN>  
 A:Cross-references: GB:X00673; NID:9192; PIDN:CAA2586.1; PID:9193  
 R:Enfield, D.L.; Ericsson, L.H.; Fujikawa, K.; Walsh, K.A.; Neurath, H.; Titani, K.  
 Biochemistry 19, 659-667, 1980  
 A:Title: Amino acid sequence of the light chain of bovine factor X-1 (Stuart factor).  
 A:Reference number: A14997; MUID:80130563; PMID:6766735  
 A:Accession: A14997  
 A:Molecule type: protein  
 A:Residues: 41-102, N, 104-180 <ENF>  
 R:McMullen, B.A.; Fujikawa, K.; Kistiel, W.  
 Biochem. Biophys. Res. Commun. 115, 8-14, 1983  
 A:Title: The occurrence of beta-hydroxyaspartic acid in the vitamin K-dependent blood co  
 A:Reference number: A20274; MUID:8330813; PMID:6688526  
 A:Contents: annotation: revision to residue 103  
 R:Titani, K.; Fujikawa, K.; Enfield, D.L.; Ericsson, L.H.; Walsh, K.A.; Neurath, H.  
 Proc. Natl. Acad. Sci. U.S.A. 72, 3082-3086, 1975  
 A:Title: Bovine factor X-1 (Stuart factor): amino-acid sequence of heavy chain.  
 A:Reference number: A12030; MUID:76053069; PMID:1059093  
 A:Accession: A12030  
 A:Molecule type: protein  
 A:Residues: 183-282, 294-295, 'GDE', 299-334, 336-348, 'AE', 351-354, 356-441, 'GKRG', 446-492 <T  
 A:Note: carboxylate binding sites and disulfide bonds were determined  
 R:Peterson, E.; Selaender, M.; Linse, S.; Drakenberg, T.; Czehlin, A.K.; Stenflo, J.  
 J. Biol. Chem. 264, 16897-16904, 1989  
 A:Title: Calcium binding to the isolated beta-hydroxyaspartic acid-containing epidermal  
 A:Reference number: A34412; MUID:89380326; PMID:2789221  
 A:Accession: A34412  
 A:Molecule type: protein  
 A:Residues: 85-126 <PER>  
 R:Imoue, K.; Morita, T.  
 Eur. J. Biochem. 218, 153-163, 1993  
 A:Title: Identification of O-linked oligosaccharide chains in the activation peptides of  
 A:Reference number: S39414; MUID:94062825; PMID:8243461  
 A:Accession: S39414  
 A:Molecule type: protein  
 A:Residues: 183-196, 199-209, 216-233 <INO>  
 A:Note: carboxylate binding sites  
 R:Titani, K.; Hermanson, M.A.; Fujikawa, K.; Ericsson, L.H.; Walsh, K.A.; Neurath, H.; D  
 Biochemistry 11, 4899-4903, 1972  
 A:Title: Bovine factor X-1a (activated Stuart factor). Evidence of homology with mammal  
 A:Reference number: A12453; MUID:73053314; PMID:4264286  
 A:Contents: annotation: active site  
 R:Fujikawa, K.; Titani, K.; Davie, E.W.  
 Proc. Natl. Acad. Sci. U.S.A. 72, 3359-3363, 1975  
 A:Title: Activation of bovine factor X (Stuart factor): conversion of factor Xaalpha to  
 A:Reference number: A13504; MUID:76053121; PMID:1059122  
 A:Contents: annotation: activation  
 R:Sugo, T.; Bjork, I.; Holmgren, A.; Stenflo, J.  
 J. Biol. Chem. 259, 5705-5710, 1984  
 A:Title: Calcium-binding properties of bovine factor X lacking the gamma-carboxyglutamic  
 A:Reference number: A38024; MUID:84185716; PMID:6546930  
 A:Contents: annotation: calcium binding  
 R:Morita, T.; Jackson, C.M.  
 J. Biol. Chem. 261, 4008-4014, 1986  
 A:Reference number: A38025; MUID:86140210; PMID:3949800  
 A:Contents: annotation: sulfate binding  
 C:Comment: Factor Xa converts prothrombin to thrombin during blood clotting.  
 C:Comment: The two chains are formed from a single-chain precursor by the excision of ty  
 C:Comment: The activation peptide is cleaved by factor IXa (in the intrinsic pathway), c  
 activation.  
 C:Comment: Calcium binds to the gamma-carboxyglutamic acid (Gla) residues and, with stru  
 C:Comment: The gamma-carboxyglutamic acid residues arise by a posttranslational, vitamin  
 C:Genetics:  
 A:Gene: F10  
 A:Map position: 13q34

C:Function:  
 A:Description: catalyzes the proteolytic activation of prothrombin to thrombin in the  
 A:Pathway: blood coagulation  
 C:Superfamily: coagulation factor X; EGF homology; Gla domain homology; trypsin homol  
 C:Keywords: beta-hydroxyaspartic acid; blood coagulation; calcium binding; carboxyglu  
 F:1-15/Domain: signal sequence #status predicted <PRO>  
 F:16-40/Domain: propeptide #status predicted <PRO>  
 F:41-180/Product: coagulation factor X light chain #status experimental <LCH>  
 F:90-121/Domain: EGF homology <EG1>  
 F:129-164/Domain: EGF homology <EG2>  
 F:183-492/Product: coagulation factor X heavy chain #status experimental <HCH>  
 F:183-233/Domain: activation peptide #status experimental <APT>  
 F:234-492/Product: coagulation factor Xa heavy chain #status experimental <AHC>  
 F:234-461/Domain: trypsin homology <TRY>  
 F:46-47, 54, 56, 59, 60, 65, 66, 69, 72, 75, 79/Modified site: gamma-carboxyglutamic acid (Glu)  
 F:57-62, 90-101, 95-110, 112-121, 129-140, 136-149, 151-164, 172-341/Disulfide bonds: #statu  
 F:103/Modified site: erythro-beta-hydroxyaspartic acid (Asp) #status experimental  
 F:200/Binding site: sulfate (Tyr) (covalent) (partial) #status experimental  
 F:208/485/Binding site: carboxylate (Thr) (covalent) #status experimental  
 F:218/Binding site: carboxylate (Asn) (covalent) #status experimental  
 F:233-234/Cleavage site: Arg-1le (coagulation factor IXa, coagulation factor VIIa) #s  
 F:240-245, 260-276, 389-403, 414-442/Disulfide bonds: #status experimental  
 F:275, 321, 418/Active site: His, Asp, Ser #status predicted  

Query Match	57.9%	Score 114	DB 1	Length 492
Best Local Similarity	45.5%	Pred. No. 2.5e-10		
Matches 20	Conservative	8	Mismatches 16	Indels 0
			Gaps 0	

QY 1 ANSFLLXLRGSLRXRCXICDFFXXAKKIFEDVDTLAFMKH 44  
 DB 41 ANSFLEEVKQGNLRECEACSLSEAEVFEADQTFDFMSKY 84  
 EXHU  
 coagulation factor Xa (EC 3.4.21.6) precursor [validated] - human  
 N:Alternate names: Stuart factor  
 C:Species: Homo sapiens (man)  
 C:Date: 15-Nov-1994 #sequence, revision 02-May-1994 #text, change 08-Dec-2000  
 C:Accession: A24478; J00917; A42485; A25853; A22208; A21284; A20362; S39415; I54051;  
 R:Leytus, S.P.; Foster, D.C.; Kurauchi, K.; Davie, E.W.  
 Biochemistry 25, 5098-5102, 1986  
 A:Title: Gene for human Factor X: a blood coagulation factor whose gene organization  
 A:Reference number: A24478; MUID:87026600; PMID:3768336  
 A:Accession: A24478  
 A:Molecule type: DNA  
 A:Residues: 1-488 <LEV>  
 A:Cross-references: GB:I29433; GB:M14327; NID:9459809; PIDN:AAA52764.1; PID:9182831  
 R:Meslier, T.L.; Pittman, D.D.; Long, G.L.; Kautman, R.J.; Church, W.R.  
 Gene 99, 291-294, 1991  
 A:Title: Cloning and expression in COS-1 cells of a full-length cDNA encoding human c  
 A:Reference number: J00917; MUID:91216473; PMID:1902434  
 A:Accession: J00917  
 A:Molecule type: mRNA  
 A:Residues: 1-488 <MES>  
 A:Cross-references: GB:M57285; NID:9182389; PIDN:AAA54221.1; PID:9182390  
 R:Miao, C.H.; Leytus, S.P.; Chung, S.P.; Davie, E.W.  
 J. Biol. Chem. 267, 7395-7401, 1992  
 A:Title: Liver-specific expression of the gene coding for human factor X, a blood coa  
 A:Reference number: A42485; MUID:92218390; PMID:1313796  
 A:Accession: A42485  
 A:Molecule type: DNA  
 A:Residues: 1-15 <MIA>  
 A:Experimental source: liver  
 A:Note: sequence extracted from NCI backbone (NCBI:93780, NCBIP:93787)  
 R:Kaul, R.K.; Hildebrand, B.; Roberts, S.; Jagadeeswaran, P.  
 Gene 41, 311-314, 1986  
 A:Title: Isolation and characterization of human blood-coagulation factor X cDNA.  
 A:Reference number: A25853; MUID:86221713; PMID:3011603  
 A:Accession: A25853  
 A:Molecule type: mRNA  
 A:Residues: 19-284, 'E', 289-488 <KAU>





A:Residues: 61-212 <THI>  
 A:Accession: B31166  
 A:Molecule type: protein  
 A:Residues: 213-466 <TH2>  
 R:Bojarn, S.; Foster, D.C.; Thim, L.; Wiberg, F.C.; Christensen, M.; Komiyama, Y.; Peder J. Biol. Chem. 266, 11051-11057, 1991  
 A:Title: Human plasma and recombinant factor VII. Characterization of O-glycosylations  
 A:Reference number: A40529; MUID:91250411; PMID:1904059  
 A:Contents: annotation; carbohydrate binding sites  
 R:Persson, E.; Petersen, L.C.  
 Eur. J. Biochem. 234, 293-300, 1995  
 A:Title: Structurally and functionally distinct Ca(2+) binding sites in the gamma-carbox  
 A:Reference number: 563524; MUID:96096552; PMID:852655  
 A:Accession: 563524  
 A:Molecule type: protein  
 A:Residues: 61-65;99-103;105-109;213-217;308-312 <PER>  
 A:Genetics:  
 A:Gene: GDB:F7  
 A:Cross-references: GDB:119897; OMIM:227500  
 A:Map position: 13q34-13q34  
 A:Introns: 22/1; 44/1; 97/3; 106/1; 144/1; 191/1; 227/3; 269/1  
 C:Function:  
 A:Description: catalyzes the proteolytic activation of coagulation factor X in the prese  
 coagulation factor IX in the presence of calcium and tissue factor  
 A:Pathway: blood coagulation extrinsic pathway  
 C:Superfamily: coagulation factor X; EGF homology; Gla domain homology; trypsin homology  
 C:Keywords: beta-hydroxyaspartic acid; blood coagulation; calcium binding; carboxylglut  
 F:1-20/Domains: signal sequence #status predicted <SIG>  
 F:21-60/Domains: propeptide #status predicted <PRO>  
 F:45-104/Domains: Gla domain homology <Gla>  
 F:61-212/Product: coagulation factor VIIa light chain #status experimental <MA1>  
 F:110-141/Domains: EGF homology <EG1>  
 F:151-187/Domains: EGF homology <EG2>  
 F:213-466/Product: coagulation factor VIIa heavy chain #status experimental <MA2>  
 F:213-447/Domains: trypsin homology <TRY>  
 F:66;67;74;76;79;85;86;89;95/Modified site: gamma-carboxylglutamic acid (Glu) #status  
 F:77-82;110-121;115-130;132-141;151-162;158-172;174-187;195-322;219-224;238-254;370-389,  
 F:112;120/Binding site: carboxylate (Ser) (covalent) #status experimental  
 F:132/Modified site: erythro-beta-hydroxyaspartic acid (Asp) #status absent  
 F:205;382/Binding site: carboxylate (Asn) (covalent) #status experimental  
 F:212-213/Cleavage site: Arg-116 (coagulation factor XIa) #status experimental  
 F:251;302;404/Active site: His, Asp, Ser #status predicted  
 F:350-351/Cleavage site: Arg-Gly (coagulation factor Xa) #status predicted

Query Match 50.3%; Score 99; DB 1; Length 466;  
 Best Local Similarity 48.8%; Pred. No. 6;Le-08;  
 Matches 20; Conservative 4; Mismatches 17; Indels 0; Gaps 0;

OY 1 ANSFLXLRGSLKRCIXXICDPXAKXIFEDVDLAFW 41  
 Db 61 ANAFLELRPGSLERCKEEOCSFEAEPRFKDAERTKLEFW 101

RESULT 10  
 S10511  
 thrombin (EC 3.4.21.5) precursor - rat  
 C:Species: Rattus norvegicus (Norway rat)  
 C:Date: 07-May-1993 #sequence,revision 07-May-1993 #text,change 03-May-2002  
 C:Accession: S10511; A60576; B42696  
 R:Dhanich, M.; Monard, D.  
 Nucleic Acids Res. 18, 4251, 1990  
 A:Title: cDNA sequence of rat prothrombin.  
 A:Reference number: S10511; MUID:90332426; PMID:2377469  
 A:Accession: S10511  
 A:Molecule type: mRNA  
 A:Residues: 1-617 <DTH>  
 A:Cross-references: EMBL:X52835; NID:956969; PIDD:CA37017.1; PID:956970  
 R:Henrikson, K.P.; Jazin, E.E.; Greenwood, J.A.; Dickerman, H.W.  
 Endocrinology 126, 167-175, 1990  
 A:Title: Prothrombin levels are increased in the estrogen-treated immature rat uterus.  
 A:Reference number: A60576; MUID:90091942; PMID:2293980  
 A:Accession: A60576  
 A:Molecule type: protein

A:Residues: 44-58 <HEN>  
 A>Note: the authors purified the proenzyme from the estrogen-stimulated maturing rat  
 R:Banfield, D.K.; MacGillivray, R.T.A.  
 Proc. Natl. Acad. Sci. U.S.A. 89, 2779-2783, 1992  
 A:Title: Partial characterization of vertebrate prothrombin cDNAs: amplification and  
 A:Reference number: A42696; MUID:92212913; PMID:1557383  
 A:Accession: B42696  
 A>Status: preliminary  
 A:Molecule type: mRNA  
 A:Residues: 383-617, 'E' <BAN>  
 A:Cross-references: GB:M81397  
 C:Superfamily: thrombin; Gla domain homology; kringle homology; trypsin homology  
 C:Keywords: blood coagulation; calcium binding; carboxylglutamic acid; glycoprotein; h  
 F:1-24/Domains: signal sequence #status predicted <SIG>  
 F:25-43/Domains: propeptide #status predicted <PRO>  
 F:28-88/Domains: Gla domain homology <Gla>  
 F:44-617/Product: prothrombin #status experimental <PMAT>  
 F:109-187/Domains: kringle homology <KR1>  
 F:215-293/Domains: kringle homology <KR2>  
 F:215-292/Domains: kringle homology <KR2>  
 F:360-609/Domains: trypsin homology <TRY>  
 F:50;51;58;60;63;64;69;70;73;76/Modified site: gamma-carboxylglutamic acid (Glu) #stat  
 F:61-66;91-104;109-187;130-170;158-182;215-292;236-276;264-287;332-478;387-403;532-54  
 F:402;458;564/Active site: His, Asp, Ser #status predicted

Query Match 43.9%; Score 86.5; DB 2; Length 617;  
 Best Local Similarity 42.2%; Pred. No. 8;Se-06;  
 Matches 19; Conservative 6; Mismatches 19; Indels 1; Gaps 1;

OY 1 ANSFLXLRGSLKRCIXXICDPXAKXIFEDVDLAFW 44  
 Db 44 ANSFLLELRKGNLERCKEEOCSFEAEPRFKDAERTKLEFW 88

RESULT 11  
 A35827  
 thrombin (EC 3.4.21.5) precursor - mouse  
 C:Species: Mus musculus (house mouse)  
 C:Date: 14-Dec-1990 #sequence,revision 14-Dec-1990 #text,change 03-May-2002  
 C:Accession: A35827; A42696; S12081  
 R:Deegen, S.J.F.; Schaefer, L.A.; Jamison, C.S.; Grant, S.G.; Fitzgibbon, J.J.; Pal, J  
 DNA Cell Biol. 9, 487-498, 1990  
 A:Title: Characterization of the cDNA coding for mouse prothrombin and localization o  
 A:Reference number: A35827; MUID:91025551; PMID:2222810  
 A:Accession: A35827  
 A>Status: preliminary  
 A:Molecule type: mRNA  
 A:Residues: 1-618 <DEG>  
 A:Cross-references: GB:X52308; NID:953813; PIDD:CA36548.1; PID:953814  
 A:Experimental source: Strain C57BL/6  
 A>Note: The data were obtained from females resulting from the cross of M. domesticus  
 R:Banfield, D.K.; MacGillivray, R.T.A.  
 Proc. Natl. Acad. Sci. U.S.A. 89, 2779-2783, 1992  
 A:Title: Partial characterization of vertebrate prothrombin cDNAs: amplification and  
 A:Reference number: A42696; MUID:92212913; PMID:1557383  
 A:Accession: A42696  
 A>Status: preliminary  
 A:Molecule type: mRNA  
 A:Residues: 384-618, 'E' <BAN>  
 A:Cross-references: GB:M81394  
 C:Superfamily: thrombin; Gla domain homology; kringle homology; trypsin homology  
 C:Keywords: blood coagulation; calcium binding; carboxylglutamic acid; glycoprotein; h  
 F:1-24/Domains: signal sequence #status predicted <SIG>  
 F:25-43/Domains: propeptide #status predicted <PRO>  
 F:28-88/Domains: Gla domain homology <Gla>  
 F:44-618/Product: prothrombin B #status predicted <MAT>  
 F:109-187/Domains: kringle homology <KR1>  
 F:215-293/Domains: kringle homology <KR2>  
 F:361-610/Domains: trypsin homology <TRY>  
 F:50;51;58;60;63;64;69;70;73;76/Modified site: gamma-carboxylglutamic acid (Glu) #stat  
 F:61-66;91-104;109-187;130-170;158-182;215-293;236-276;264-288;333-479;388-404;533-54  
 F:403;459;565/Active site: His, Asp, Ser #status predicted

Query Match 43.9%; Score 86.5; DB 2; Length 618;



Best Local Similarity 42.2%, Pred. No. 8.5e-06;  
Matches 19; Conservative 6; Mismatches 19; Indels 1; Gaps 1;  
Oy 1 ANS-FLXXLRGSLKRXICIXXICDFXXAKXIFEDVDLTAFLWS 44  
Db 44 ANSGFLEELKRGKGNLERECVEECSTEEAFEALESPODIDVETAKY 88

## RESULT 12

## EXCH

coagulation factor Xa (EC 3.4.21.6) precursor - chicken  
N:Alternate names: virus-activating proteinase

C:Species: Gallus gallus (chicken)

C:Date: 12-Feb-1993 #sequence\_revision 07-Feb-1997 #text\_change 16-Jul-1999

C:Accession: S15838; S20380; S20381

R:Stuuk, H.; Harada, A.; Hayashi, Y.; Wada, K.; Asaka, J.; Gotoh, B.; Ogasawara, T.; NE  
FEBS Lett. 283: 281-285, 1991

A:Title: Primary structure of the virus activating protease from chick embryo. Its ident

A:Reference number: S15838; MUID:91257322; PMID:2044767

A:Accession: S15838

A:Status: not compared with conceptual translation

A:Molecule type: mRNA

A:Residues: 1-475 <SUZ>

A:Cross-references: DDBJ:D00844; NID:9222869; PIDN:BA00724.1; PID:9222870

R:Gotoh, B.; Yamauchi, F.; Ogasawara, T.; Nagai, Y.  
FEBS Lett. 296: 274-278, 1992

A:Title: Isolation of factor Xa from chick embryo as the amniotic endoprotease responsib

A:Reference number: S20380; MUID:92164779; PMID:1537403

A:Accession: S20380

A:Molecule type: protein

A:Residues: 41-55 <GO2>

A:Accession: S20381

A:Molecule type: protein

A:Residues: 241-246, 'X', 248-251, 'X', 253-261 <GOT>

C:Function:

A:Description: catalyzes the proteolytic activation of prothrombin to thrombin in the pr

A:Pathway: blood coagulation

C:Superfamily: coagulation factor X; EGF homology; Gla domain homology; trypsin homology

C:Keywords: beta-hydroxyaspartic acid; blood coagulation; calcium binding; carboxyglutan

F:1-20/Domain: signal sequence #status predicted <SIG>

F:21-40/Domain: propeptide #status predicted <PRO>

F:41-185/Product: coagulation factor X light chain #status experimental <LCH>

F:90-121/Domain: EGF homology <EG1>

F:129-167/Domain: EGF homology <EG2>

F:186-475/Product: coagulation factor X heavy chain #status predicted <HCH>

F:186-240/Domain: activation peptide #status predicted <AP>

F:241-475/Product: coagulation factor Xa heavy chain #status experimental <AHCH>

F:241-468/Domain: trypsin homology <TRY>

F:46-47, 54, 56, 59, 60, 65, 66, 69, 72, 75, 79/Modified site: gamma-carboxyglutamic acid (Glu) #

F:57-62, 90-101, 95-110, 112-121, 129-140, 136-152, 154-167, 175-348, 247-252, 267-283, 396-410, 42

F:103/Modified site: erythro-beta-hydroxyaspartic acid (Asp) #status predicted

F:196, 207, 228, 285/Binding site: carbonyl (Asn) (covalent) #status predicted

F:283, 328, 425/Active site: His, Asp, Ser #status predicted

Query Match 43.7%, Score 86; DB 1; Length 475;  
Best Local Similarity 40.5%; Pred. No. 7.8e-06;  
Matches 17; Conservative 7; Mismatches 18; Indels 0; Gaps 0;

Oy 1 ANSFLXLRGSLKRXICIXXICDFXXAKXIFEDVDLTAFLWS 42  
Db 41 ANSFLEEMKOGNIEECNEERCSKEAREAFEDNEKEEFVNW 82

## RESULT 13

## KFB07

coagulation factor VIIa (EC 3.4.21.21) - bovine

C:Species: Bos primigenius taurus (cattle)

C:Date: 21-May-1990 #sequence\_revision 23-Mar-1995 #text\_change 16-Jul-1999

C:Accession: A31979; C20274

R:Takeya, H.; Kawabata, S.; Nakagawa, K.; Yamamichi, Y.; Miyata, T.; Iwanaga, S.  
J. Biol. Chem. 265: 14868-14877, 1988

A:Title: Bovine factor VII. Its purification and complete amino acid sequence.

A:Reference number: A31979; MUID:89008362; PMID:3049594  
A:Accession: A31979  
A:Molecule type: protein  
A:Residues: 1-407 <TAK>

R:McMullen, B.A.; Fujikawa, K.; Kistel, W.

Biochem. Biophys. Res. Commun. 115: 8-14, 1983

A:Title: The occurrence of beta-hydroxyaspartic acid in the vitamin K-dependent blood

A:Reference number: A20274; MUID:83308813; PMID:6688526

A:Accession: C20274

A:Molecule type: protein

A:Residues: 58-62, 'X', 64-68 <MCK>

A>Note: The residue designated 'X' was determined to be hydroxyaspartic acid

R:Hase, S.; Kawabata, S.; Nishimura, H.; Takeya, H.; Sueyoshi, T.; Miyata, T.; Iwanag

J. Biochem. 104: 867-868, 1988

A:Title: A new trisaccharide sugar chain linked to a serine residue in bovine blood c

A:Reference number: A44556; MUID:89213999; PMID:3149637

A:Contents: annotation

A>Note: structure and location of covalently bound carbohydrate

C:Function:

A:Description: catalyzes the proteolytic activation of coagulation factor X in the pr

A:Pathway: blood coagulation extrinsic pathway

C:Superfamily: coagulation factor X; EGF homology; Gla domain homology; trypsin homol

C:Keywords: beta-hydroxyaspartic acid; blood coagulation; calcium binding; carboxyglu

F:1-152/Product: coagulation factor VIIa light chain #status experimental <MA1>

F:1-44/Domain: Gla domain homology (fragment) <GUA>

F:50-81/Domain: EGF homology <EG1>

F:91-127/Domain: EGF homology <EG2>

F:153-407/Product: coagulation factor VIIa heavy chain #status experimental <MA2>

F:153-387/Domain: trypsin homology <TRY>

F:6, 7, 14, 16, 19, 20, 25, 26, 29, 34, 35/Modified site: gamma-carboxyglutamic acid (Glu) #sta

F:17-22, 50-61, 55-70, 72-81, 91-102, 98-112, 114-127, 135-262, 159-164, 178-194, 310-329, 340-3

F:52/Binding site: carbonyl (Ser) (covalent) #status experimental

F:145, 203/Binding site: erythro-beta-hydroxyaspartic acid (Asp) (partial) #status experim

F:152-153/Cleavage site: Arg-Tile (coagulation factor XIIIa) #status experimental

F:193, 242, 344/Active site: His, Asp, Ser #status predicted

F:290-291/Cleavage site: Arg-Gly (coagulation factor Xa) #status experimental

Query Match 43.1%, Score 85; DB 1; Length 407;  
Best Local Similarity 43.9%; Pred. No. 9.7e-06;  
Matches 18; Conservative 3; Mismatches 20; Indels 0; Gaps 0;

Oy 1 ANSFLXLRGSLKRXICIXXICDFXXAKXIFEDVDLTAFLW 41  
Db 1 ANGFLEELKRGKGNLERECVEECSTEEAFEALESPODIDVETAKY 41

## RESULT 14

## S53434

plasma protein S precursor, vitamin K dependent - rhesus macaque (fragment)

C:Species: Macaca mulatta (rhesus macaque)

C:Date: 19-Mar-1997 #sequence\_revision 18-Jul-1997 #text\_change 16-Jul-1999

C:Accession: S53434

R:Greengard, J.S.; Fernandez, J.A.; Radtke, K.P.; Griffin, J.H.

Biochem. J. 305: 397-403, 1995

A:Title: Identification of candidate residues for interaction of protein S with C4b b

A:Reference number: S53433; MUID:95134217; PMID:7832752

A:Accession: S53434

A:Status: nucleic acid sequence not shown; not compared with conceptual translation

A:Molecule type: mRNA

A:Residues: 1-642 <GRE>

A:Cross-references: EMBL:L31380

A:Experimental source: tissue type liver

A>Note: the source is designated as rhesus monkey

C:Genetics:

A:Gene: PROS

C:Superfamily: plasma protein S; EGF homology; Gla domain homology; vitamin G repeat

F:1-51/Domain: Gla domain homology (fragment) <GLA>

F:1-7/Domain: signal sequence (fragment) #status predicted <SIG>

F:8-642/Product: plasma protein S #status predicted <MAT>

F:87-120/Domain: EGF homology <EG1>

F:127-165/Domain: EGF homology <EG2>

F:171-207/Domain: EGF homology <EG3>  
 F:213-248/Domain: EGF homology <EG4>  
 F:281-633/Domain: sex hormone-binding globulin homology <SHB>  
 F:291-444/Domain: laminin G repeat homology <LGR>

## Query Match

Best Local Similarity 43.1%; Score 85; DB 2; Length 642;  
 Pred. No. 1.5e-05;

Matches 17; Conservative 10; Mismatches 17; Indels 0; Gaps 0;  
 OY 1 ANSFLXXRGSIXRCIXXICDFXXAKXIFEDVDTLAFMSKH 44  
 DB 8 ANSMLEETKOGNLEKRECEIELCNKEAREVEFNDPETYFYPKY 51

## RESULT 15

KXKHUS

Plasma protein S precursor - human

C:Species: Homo sapiens (man)

C:Date: 21-Sep-1990 #sequence.revision 26-Jan-1996 #text.change 16-Jul-1999

C:Accession: A35610; A35611; A26157; A25891; A35612; A60903; S02424; S09519

R:Schmidel, D.K.; Tatro, A.V.; Phelps, L.G.; Tomczak, J.A.; Long, G.L.

Biochemistry 29, 7845-7852, 1990

A:Title: Organization of the human protein S genes.

A:Reference number: A35610; MUID:91084444; PMID:2148110

A:Accession: A35610

A:Molecule type: DNA

A:Residues: 1-676 &lt;SCH&gt;

A:Cross-references: GB:M57853; NID:q19054; PIDN:AAA60357.1; PID:q190549; GB:J02917

R:Note: the authors translated the codon TTT for residue 26 as Leu

R:Ploos van Amstel, H.K.; Reitsma, P.H.; van der Logt, C.P.E.; Bertina, R.M.

Biochemistry 29, 7853-7861, 1990

A:Title: Intron-exon organization of the active human protein S gene. Psalpa and its pse

A:Reference number: A35611; MUID:91084445; PMID:2148111

A:Accession: A35611

A:Molecule type: DNA

A:Residues: 1-25 &lt;PL3&gt;

A:Cross-references: GB:J02918

R:Hoskins, J.; Norman, D.K.; Beckmann, R.J.; Long, G.L.

Proc. Natl. Acad. Sci. U.S.A. 84, 349-353, 1987

A:Title: Cloning and characterization of human liver cDNA encoding a protein S precursor

A:Reference number: A26157; MUID:87092407; PMID:3467362

A:Accession: A26157

A:Molecule type: mRNA

A:Residues: 1-10, 'P', '12-25, 'L', '27-676 &lt;HOS&gt;

A:Cross-references: GB:M15036; NID:q190288; PIDN:AAA36479.1; PID:q190289

R:Lundwall, A.; Backowski, W.; Cohen, E.; Shaffer, M.; Manr, A.; Dahlback, B.; Stenflo,

Proc. Natl. Acad. Sci. U.S.A. 83, 6716-6720, 1986

A:Title: Isolation and sequence of the cDNA for human protein S, a regulator of blood co

A:Reference number: A25891; MUID:86313649; PMID:2944113

A:Accession: A25891

A:Molecule type: mRNA

A:Residues: 27-220, 'L', '222-262, 'H', '264-344, 'Y', '345-676 &lt;LUN&gt;

A:Cross-references: GB:M14338; NID:q190448; PIDN:AAA60181.1; PID:q190449

R:Benndorf, C.M.; Lundwall, A.; Wydro, R.; Stenflo, J.

Biochemistry 29, 7861-7868, 1990

A:Title: Molecular analysis of the gene for vitamin K dependent protein S and its pseud

A:Reference number: A35612; MUID:91084446; PMID:2148112

A:Accession: A35612

A:Molecule type: DNA

A:Residues: 284-676 &lt;EDR&gt;

A:Cross-references: GB:J02919

R:Ploos van Amstel, J.K.; van der Zanden, A.L.; Bakker, E.; Reitsma, P.H.; Bertina, R.M.

Thromb. Haemost. 58, 982-987, 1987

A:Title: Two genes homologous with human protein S cDNA are located on chromosome 3.

A:Reference number: A60903; MUID:88178564; PMID:2895503

A:Accession: A60903

A:Molecule type: mRNA

A:Residues: 351-676 &lt;PIO&gt;

R:Ploos van Amstel, H.K.; van der Zanden, A.L.; Reitsma, P.H.; Bertina, R.M.

FEBS Lett. 222, 186-190, 1987

A:Title: Human protein S cDNA encodes Phe-16 and Tyr 222 in consensus sequences for t  
 A:Reference number: S02424; MUID:88005138; PMID:2820795

A:Accession: S02424

A:Molecule type: mRNA

A:Residues: 1-676 &lt;PL2&gt;

A:Cross-references: EMBL:Y00692; NID:q36578; PIDN:CAA68867.1; PID:q36579

C:Gene: GDB:PROS1; PROS

A:Cross-references: GDB:120721; OMIM:176880

A:Map position: 3p11.1-3q11.2

A:Introns: 26/1; 78/3; 87/1; 116/1; 157/1; 201/1; 243/1; 283/3; 322/2; 385/3; 441/3;

C:Complex: In plasma forms a complex with C4b binding protein

C:Function:

A:Description: A cofactor for activated protein C (EC 3.4.21.69); thrombin cleavage d

C:Superfamily: plasma protein S; EGF homology; Gla domain homology; laminin G repeat

C:Keywords: beta-hydroxyasparagine; beta-hydroxyaspartic acid; blood coagulation; car

F:1-24/Domain: signal sequence #status predicted &lt;SIG&gt;

F:25-41/Domain: propeptide #status predicted &lt;PRO&gt;

F:42-67/Domain: Gla domain homology &lt;Gla&gt;

F:121-154/Domain: EGF homology &lt;EG1&gt;

F:161-199/Domain: EGF homology &lt;EG2&gt;

F:205-241/Domain: EGF homology &lt;EG3&gt;

F:247-282/Domain: EGF homology &lt;EG4&gt;

F:315-667/Domain: sex hormone-binding globulin homology &lt;SHB&gt;

F:325-478/Domain: laminin G repeat homology &lt;LGR&gt;

F:47/48, 55, 57, 60, 61, 66, 67, 70, 73, 77/Modified site: gamma-carboxyglutamic acid (Glu) #s

F:58-63, 88-113, 121-134, 126-143, 145-154, 161-175, 171-184, 186-199, 205-217, 212-226, 228-24

F:111-112/Cleavage site: Arg-Ser (thrombin) #status predicted

F:136/Modified site: erythro-beta-hydroxyaspartic acid (Asp) #status predicted

F:177, 219, 258/Modified site: erythro-beta-hydroxyasparagine (Asn) #status predicted

F:499, 509, 530/Binding site: carboxylate (Asn) (covalent) #status predicted

## Query Match

Best Local Similarity 43.1%; Score 85; DB 1; Length 676;  
 Pred. No. 1.6e-05;

Matches 17; Conservative 10; Mismatches 17; Indels 0; Gaps 0;  
 OY 1 ANSFLXXRGSIXRCIXXICDFXXAKXIFEDVDTLAFMSKH 44  
 DB 42 ANSMLEETKOGNLEKRECEIELCNKEAREVEFNDPETYFYPKY 85

Search completed: May 16, 2003, 10:15:51

Job time : 20 secs

GenCore version 5.1.4-p5.4578  
Copyright (c) 1993 - 2003 CompuGen Ltd.

OM protein - protein search, using sw model

Run on: May 16, 2003, 10:12:18 ; Search time 11 Seconds

(without alignments)  
165.905 Million cell updates/sec

Title: SEQ1-4EDITS

Perfect score: 197  
Sequence: 1 ANSFLXLRQGSIXRXCIX.....XXAKXIFEDVDDTLAFWSKH 44

Scoring table: BLOSUM62  
Gapop 10.0 , Gapext 0.5

Searched: 112892 seqs, 41476328 residues

Total number of hits satisfying chosen parameters: 112892

Minimum DB seq length: 0

Maximum DB seq length: 200000000

Post-processing: Minimum Match 0%

Maximum Match 100%

Database : SwissProt\_40.\*

Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

## SUMMARIES

Result No.	Score	Query Match	Length	ID	Description
1	160	81.2	461	1 PRTC_HUMAN	P04070 homo sapien
2	140	71.1	461	1 PRTC_MOUSE	P33587 mus musculu
3	139	70.6	461	1 PRTC_RAT	P31394 rattus norv
4	138	70.1	458	1 PRTC_RABIT	Q28661 oryctolagus
5	123	62.4	459	1 PRTC_PIG	Q99122 sus scrofa
6	122	61.9	456	1 PRTC_BOVIN	P00745 bos taurus
7	114	57.9	492	1 FA10_BOVIN	P00743 bos taurus
8	110	55.8	488	1 FA10_HUMAN	P00742 homo sapien
9	107	54.3	231	1 TMG3_HUMAN	Q9bzd7 homo sapien
10	103	52.3	490	1 FA10_RABIT	O19045 oryctolagus
11	101	51.3	444	1 FA7_RABIT	P98139 oryctolagus
12	99	50.3	466	1 FA7_HUMAN	P08709 homo sapien
13	92	46.7	218	1 TMG1_HUMAN	O14668 homo sapien
14	86.5	43.9	617	1 THRB_RAT	P18292 rattus norv
15	86.5	43.9	618	1 THRB_MOUSE	P19221 mus musculu
16	86	43.7	475	1 FA10_CHICK	P25155 gallus gall
17	85	43.1	407	1 FA7_BOVIN	P22457 bos taurus
18	85	43.1	649	1 PRTS_MACMU	Q28520 macaca mula
19	85	43.1	676	1 PRTS_HUMAN	P07725 homo sapien
20	84.5	42.6	226	1 TMG4_HUMAN	Q9bzd6 homo sapien
21	84	42.6	622	1 THRB_HUMAN	P00734 homo sapien
22	82	41.6	376	1 FA10_TROCA	P81428 tropidochis
23	81	41.1	646	1 PRTS_RABIT	P98118 oryctolagus
24	80	40.6	446	1 FA7_MOUSE	P70375 mus musculu
25	80	40.6	452	1 FA9_CANFA	P19540 canis faml
26	80	40.6	459	1 FA9_MOUSE	P16294 mus musculu
27	80	40.6	461	1 FA9_HUMAN	P00740 homo sapien
28	80	40.6	675	1 PRTS_BOVIN	P07224 bos taurus
29	78	39.6	675	1 PRTS_RAT	P51813 rattus norv
30	73	37.1	416	1 FA9_BOVIN	P00741 bos taurus
31	72	36.5	625	1 THRB_BOVIN	P00735 bos taurus
32	71	36.0	675	1 PRTS_MOUSE	Q08761 mus musculu
33	69.5	35.3	396	1 PRT2_BOVIN	P00744 bos taurus

34	65.5	33.2	400	1 PRT2_HUMAN	P22891 homo sapien
35	65	33.0	202	1 TMG2_HUMAN	O14669 homo sapien
36	52	26.4	501	1 MKC1_CANAL	P43068 candida alb
37	50	25.4	363	1 ADK_TOXGO	Q91vw2 toxoplasma
38	49	24.9	1363	1 VGR3_MOUSE	P35917 mus musculu
39	48	24.4	422	1 SPML_SCHPO	Q92398 schizosacch
40	48	24.4	1235	1 CYA4_TRYBB	Q26721 trypanosoma
41	48	24.4	1298	1 VGR3_HUMAN	P35916 homo sapien
42	47	23.9	244	1 T2E5_ECOLI	P04390 escherichia
43	47	23.9	554	1 DHAE_SALTY	P37450 salmonella
44	47	23.9	1343	1 VGR2_RAT	O08775 rattus norv
45	47	23.9	1348	1 VGR2_COTJA	P52583 coturnix co

## ALIGNMENTS

RESULT 1  
PRTC\_HUMAN STANDARD: PRT; 461 AA.  
ID AC P04070; 016001; 015190; 015189;  
DT 01-NOV-1986 (Rel. 03, Last sequence update)  
DT 01-NOV-1986 (Rel. 03, Last sequence update)  
DE 15-JUN-2002 (Rel. 41, Last annotation update)  
DE Vitamin-K dependent protein C precursor (EC 3.4.21.69)  
DE (Autoproteolytic IIA) (Anticoagulant protein C) (Blood coagulation factor XIV).  
GN PROC.  
OS Homo sapiens (Human).  
OC Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
OC Mammalia; Eutheria; Primates; Catarrhini; Homiidae; Homo.  
OX NCBI\_TaxID=9606;  
RN [1]  
RP SEQUENCE FROM N.A.  
RX MEDLINE=85270390; PubMed=2991887;  
RA Foster D.C., Yoshitake S., Davie E.W.;  
RT "The nucleotide sequence of the gene for human protein C";  
RL Proc. Natl. Acad. Sci. U.S.A. 82:4673-4677(1985).  
RN [2]  
RP SEQUENCE FROM N.A.  
RX MEDLINE=85269639; PubMed=2991859;  
RA Beckmann R.J., Schmidt R.J., Santerre R.F., Plutsky J., Crabtree G.R.,  
RN Long G.L.;  
RT "The structure and evolution of a 461 amino acid human protein C  
RT precursor and its messenger RNA, based upon the DNA sequence of  
RT cloned human liver cDNAs";  
RL Nucleic Acids Res. 13:5233-5247(1985).  
RN [3]  
RP SEQUENCE FROM N.A.  
RX MEDLINE=86120978; PubMed=3511471;  
RA Plutsky J., Hoskins J.A., Long G.L., Crabtree G.R.;  
RT "Evolution and organization of the human protein C gene";  
RL Proc. Natl. Acad. Sci. U.S.A. 83:546-550(1986).  
RN [4]  
RP SEQUENCE FROM N.A.  
RX Rieder M.J., Carrington D.P., Chung M.-W., Lee K.L., Peol C.L., Yi Q.,  
RN Nickerson D.A.;  
RT Submitted (JUN-2001) to the EMBL/GenBank/DBJ databases.  
RN [5]  
RP SEQUENCE OF 106-461 FROM N.A.  
RX MEDLINE=84272714; PubMed=6589623;  
RA Foster D.C., Davie E.W.;  
RT "Characterization of a cDNA coding for human protein C";  
RL Proc. Natl. Acad. Sci. U.S.A. 81:4766-4770(1984).  
RN [6]  
RP CARBOHYDRATE-LINKAGE SITE ASN-371.  
RX MEDLINE=90293094; PubMed=1694179;  
RA Mjelich J.P., Broze G.J. Jr.;  
RT "Beta protein C is not glycosylated at asparagine 329. The rate of  
RT translation may influence the frequency of usage at asparagine-X-  
RT cysteine sites";  
RL J. Biol. Chem. 265:11397-11404(1990).  
RN [7]

RP HYDROXYLATION.  
RX MEDLINE-92184750; PubMed-1544894;  
RA Harris R.J., Ling V.T., Spellman M.W.;  
RT "O-linked fucose is present in the first epidermal growth factor  
RL domain of factor XII but not protein C.";  
RN J. Biol. Chem. 267:5102-5107(1992).  
RP [8]  
RP 3D-STRUCTURE MODELING OF 175-450.  
RX MEDLINE-94272342; PubMed-8003977;  
RA Fisher C.L., Greengard J.S., Griffin J.H.;  
RT "Models of the serine protease domain of the human antithrombotic  
RL plasma factor activated protein C and its zymogen.";  
RN Protein Sci. 3:588-599(1994).  
RP [9]  
RP X-RAY CRYSTALLOGRAPHY (2.8 ANGSTROMS) OF 84-461.  
RX MEDLINE-91157472; PubMed-9003757;  
RA Mather T., Oganessyan V., Hof P., Huber R., Foundling S., Esmon C.,  
RA Bode W.;  
RT "The 2.8 A crystal structure of Gla-domainless activated protein C.";  
RL EMBO J. 15:6822-6831(1996).  
RN [10]  
RN REVIEW ON PROC VARIANTS.  
RX MEDLINE-93190290; PubMed-8446940;  
RA Beltsma P.H., Poort S.R., Bernardi F., Gandrille S., Long G.L.,  
RA Sala N., Cooper D.N.;  
RT "Protein C deficiency: a database of mutations. For the Protein C & S  
RL Subcommittee of the Scientific and Standardization Committee of the  
RN International Society on Thrombosis and Haemostasis.";  
RN Thromb. Haemost. 69:77-84(1993).  
RN [11]  
RN VARIANT CYS-444.  
RX MEDLINE-87204221; PubMed-2437584;  
RA Romeo G., Hassan H.J., Staempfli S., Roncuazzi L., Cianetti L.,  
RA Leonardi A., Vicente V., Mannucci P.M., Bertina R.M., Peschle C.,  
RA Cortese R.;  
RT "Hereditary thrombophilia: identification of nonsense and missense  
RL mutations in the protein C gene.";  
RN Proc. Natl. Acad. Sci. U.S.A. 84:2829-2832(1987).  
RN [12]  
RN VARIANT TRP-211 (LONDON-1).  
RX MEDLINE-90098906; PubMed-2602169;  
RA Grundy C.B., Chittolte A., Talbot S., Bevan D., Kakkar V.V.,  
RA Cooper D.N.;  
RT "Protein C London 1: recurrent mutation at Arg-169 (CGG->TGG) in  
RL the protein C gene causing thrombosis.";  
RN Nucleic Acids Res. 17:10513-10513(1989).  
RN [13]  
RN VARIANT CYS-272.  
RX MEDLINE-91329836; PubMed-1868249;  
RA Beltsma P.H., Poort S.R., Allaart C.F., Briet E., Bertina R.M.;  
RT "The spectrum of genetic defects in a panel of 40 Dutch families with  
RL symptomatic protein C deficiency type I: heterogeneity and founder  
RN effects.";  
RN Blood 78:890-894(1991).  
RN [14]  
RN VARIANTS ALA-62 (VERMONT-1) AND MET-76.  
RX MEDLINE-92190481; PubMed-1347706;  
RA Bovill E.G., Tomczak J.A., Grant B., Bhushan F., Pillemer E.,  
RA Rainville I.R., Long G.L.;  
RT "Protein C Vermont: symptomatic type II protein C deficiency  
RL associated with two Gla domain mutations.";  
RN Blood 79:1456-1465(1992).  
RN [15]  
RN VARIANT ASP-418 (HONG KONG-2).  
RX MEDLINE-92305321; PubMed-1611081;  
RA Sugahara Y., Miura O., Yuen P., Aoki N.;  
RT "Protein C deficiency Hong Kong 1 and 2: hereditary protein C  
RL deficiency caused by two mutant alleles, a 5-nucleotide deletion and  
RN a missense mutation.";  
RN Blood 80:126-133(1992).  
RN [16]  
RN VARIANT LEU-289.  
RX MEDLINE-92380660; PubMed-1511988;  
RA Grundy C.B., Chisholm M., Kakkar V.V., Cooper D.N.;  
RT "A novel homozygous missense mutation in the protein C (PROC) gene  
RL causing recurrent venous thrombosis.";  
RN Hum. Genet. 89:683-684(1992).  
RN [17]  
RN VARIANTS GLN-220 AND TRP-220.  
RX MEDLINE-92380661; PubMed-1511989;  
RA Grundy C.B., Schulman S., Tengborn L., Kakkar V.V., Cooper D.N.;  
RT "Two different missense mutations at Arg 178 of the protein C (PROC)  
RL gene causing recurrent venous thrombosis.";  
RN Hum. Genet. 89:685-686(1992).  
RN [18]  
RN VARIANT GLN-220.  
RX MEDLINE-93250852; PubMed-1301959;  
RA Gandrille S., Vidaud M., Alach M., Alhenc-Gelas M., Fischer A.M.,  
RA Gouault-Heilman M., Toulon P., Flessinger J.N., Goossens M.;  
RT "Two novel mutations responsible for hereditary type I protein C  
RL deficiency: characterization by denaturing gradient gel  
RN electrophoresis.";  
RN Hum. Mutat. 1:491-500(1992).  
RN [19]  
RN VARIANT SER-334.  
RX MEDLINE-92276939; PubMed-1593215;  
RA Yamamoto K., Matsushita T., Sugura I., Takamatsu J., Iwasaki E.,  
RA Wada H., Deguchi K., Shirakawa S., Saito H.;  
RT "Homozygous protein C deficiency: identification of a novel missense  
RL mutation that causes impaired secretion of the mutant protein C.";  
RN J. Lab. Clin. Med. 119:682-689(1992).  
RN [20]  
RN VARIANTS TRP-38; CYS-42; HIS-42; GLN-271 AND ASN-294.  
RX MEDLINE-93313192; PubMed-8324221.  
RA Gandrille S., Alhenc-Gelas M., Gaussem P., Allaud M.-F., Dupuy E.,  
RA Juhan-Vague I., Alach M.;  
RT "Five novel mutations located in exons III and IX of the protein C  
RL gene in patients presenting with defective protein C anticoagulant  
RN activity.";  
RN Blood 82:159-166(1993).  
RN [21]  
RN VARIANTS G-14; Q-211; Y-244; Q-253; L-321; C-328; I-385; T-388 AND  
RN V-388.  
RX MEDLINE-93271391; PubMed-8499565;  
RA Poort S.R., Pabinger-Fasching I., Mannhalter C., Beltsma P.H.,  
RA Bertina R.M.;  
RT "Twelve novel and two recurrent mutations in 14 Austrian families  
RL with hereditary protein C deficiency.";  
RN Blood Coagul. Fibrinolysis 4:273-280(1993).  
RN [22]  
RN VARIANT TRP-57.  
RX MEDLINE-93271396; PubMed-8499568;  
RA Millar D.S., Grundy C.B., Bignelli P., Mofat E.H., Martin R.,  
RA Kakkar V.V., Cooper D.N.;  
RT "A Gla domain mutation (Arg 15->Trp) in the protein C (PROC) gene  
RL causing type 2 protein C deficiency and recurrent venous  
RN thrombosis.";  
RN Blood Coagul. Fibrinolysis 4:345-347(1993).  
RN [23]  
RN VARIANTS R-145; L-210; W-211; T-243; L-321; M-340 AND Y-426.  
RX MEDLINE-94122329; PubMed-8292730;  
RA Tsay W., Greengard J.S., Montgomery R.R., McPherson R.A., Fucci J.C.,  
RA Koester M.A., Coughlin J., Griffin J.H.;  
RT "Genetic mutations in ten unrelated American patients with  
RL symptomatic type I protein C deficiency.";  
RN Blood Coagul. Fibrinolysis 4:791-796(1993).  
RN [24]  
RN VARIANT SER-423.  
RX MEDLINE-94001606; PubMed-8398832;  
RA Marchetti G., Patrocchini P., Gemmati D., Castaman G., Rodeghiero F.,  
RA Wacey A., Cooper D.N., Tuddenham E.G., Bernardi F.;  
RT "Symptomatic type II protein C deficiency caused by a missense  
RL mutation (Gly 381->Ser) in the substrate-binding pocket.";  
RN Br. J. Haematol. 84:285-289(1993).  
RN [25]  
RN SEQUENCE OF 43-64 FROM N.A., AND VARIANT GLY-57 (YONAGO).



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DT 01-JUL-1993 (Rel. 26, Created)
DT 01-JUL-1993 (Rel. 26, Last sequence update)
DT 15-JUN-2002 (Rel. 41, Last annotation update)
DE Vitamin-K dependent protein C precursor (EC 3.4.21.69)
DE (Antithrombin IIIa) (Anticoagulant protein C) (Blood coagulation factor XIV).
GN PROC.
OS Rattus norvegicus (Rat).
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RA STRAIN-Mistart; TISSUE-Liver;
RX MEDLINE-92329550; PubMed-1627650.
RA Okafuji T., Maekawa K., Nawa K., Marumoto Y.;
RT "The cDNA cloning and mRNA expression of rat protein C.";
RL Biochim. Biophys. Acta 1131:329-332(1992).
CC -1- FUNCTION: PROTEIN C IS A VITAMIN K-DEPENDENT SERINE PROTEASE THAT
CC REGULATES BLOOD COAGULATION BY INACTIVATING FACTORS VA AND VIIIA
CC IN THE PRESENCE OF CALCIUM IONS AND PHOSPHOLIPIDS.
CC -1- CATALYTIC ACTIVITY: Degradation of blood coagulation factors Va
CC and VIIa.
CC -1- SUBUNIT: SYNTHESIZED AS A SINGLE CHAIN PRECURSOR, WHICH IS CLEAVED
CC INTO A LIGHT CHAIN AND A HEAVY CHAIN HELD TOGETHER BY A DISULFIDE
CC BOND. THE ENZYME IS THEN ACTIVATED BY THROMBIN, WHICH CLEAVES A
CC TETRADECAPEPTIDE FROM THE AMINO END OF THE HEAVY CHAIN; THIS
CC REACTION, WHICH OCCURS AT THE SURFACE OF ENDOTHELIAL CELLS, IS
CC STRONGLY PROMOTED BY THROMBOMODULIN.
CC -1- TISSUE SPECIFICITY: PLASMA; SYNTHESIZED IN THE LIVER.
CC -1- PTM: THE VITAMIN K-DEPENDENT, ENZYMAIC CARBOXYLATION OF SOME
CC GLU RESIDUES ALLOWS THE MODIFIED PROTEIN TO BIND CALCIUM.
CC -1- MISCELLANEOUS: CALCIUM ALSO BINDS, WITH STRONGER AFFINITY TO
CC ANOTHER SITE, BEYOND THE GLA DOMAIN. THIS GLA-INDEPENDENT BINDING
CC SITE IS NECESSARY FOR THE RECOGNITION OF THE
CC THROMBIN-THROMBOMODULIN COMPLEX.
CC -1- SIMILARITY: BELONGS TO PEPTIDASE FAMILY S1.
CC -1- SIMILARITY: CONTAINS 2 EGF-LIKE DOMAINS.
CC -----
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CC -----
DR EMBL: X64336; CAA45617.1; -.
DR PIR: S18994; S18994.
DR PIR: S24312; S24312.
DR HSSP: P04070; 1PCU.
DR MEROPS: S01.218; -.
DR InterPro: IPR000152; Asx_hydroxyl.
DR InterPro: IPR001314; Chymotrypsin.
DR InterPro: IPR000561; EGF-like.
DR InterPro: IPR001881; EGF_Ca.
DR InterPro: IPR002383; GLA_blood.
DR InterPro: IPR001254; Ser_protease_Try.
DR InterPro: IPR000294; VitK_dep_GLA.
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DR Pfam: PF00089; trypsin_1.
DR Pfam: PF00594; gla_1.
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FT CHAIN 199 461 PROTEIN C HEAVY CHAIN (BY SIMILARITY).
FT PEPTIDE 199 212 ACTIVATION PEPTIDE (BY SIMILARITY).
FT SITE 212 213 CLEAVAGE (BY THROMBIN) (BY SIMILARITY).
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FT DOMAIN 135 175 EGF-Like 2.
FT DOMAIN 213 461 SERINE PROTEASE.
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FT MOD_RES 55 55 GAMMA-CARBOXYGLUTAMIC ACID
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FT CARBOHYD 215 215 N-LINKED (GLCNAC. . .) (POTENTIAL).
FT CARBOHYD 291 291 N-LINKED (GLCNAC. . .) (POTENTIAL).
FT CARBOHYD 355 355 N-LINKED (GLCNAC. . .) (POTENTIAL).
SQ SEQUENCE 461 AA: 51912 MW: 846CF93664EDACDS CRC64:

Query Match 70.6%; Score 139; DB 1; Length 461;
Best Local Similarity 59.1%; Pred. No. 3.9e-16;
Matches 26; Conservative 7; Mismatches 11; Indels 0; Gaps 0;

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DB 42 ANSFLVEVRAGSLERECMEICDFEABOEIFONVEDTLAFWIKY 85

RESULT 4
PRTC_RABIT STANDARD; PRT; 458 AA.
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DT 15-DEC-1998 (Rel. 37, Last sequence update)
DT 15-JUN-2002 (Rel. 41, Last annotation update)
DE Vitamin-K dependent protein C precursor (EC 3.4.21.69)
DE (Autoproteolysin IIA) (Anticoagulant protein C) (Blood coagulation
DE factor XIV) (Fragment).
GN PROC.
OS Oryctolagus cuniculus (Rabbit).
OC Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC Mammalia; Eutheria; Lagomorpha; Leporidae; Oryctolagus.
OX NCBI_Taxid-9986;
RN 11)
RP SEQUENCE FROM N.A.
RC TISSUE=Liver;
RA Shen L., He X., Dahlback B.;
RL Submitted (FEB-1996) to the EMBL/Genbank/DDI databases.
CC -1- FUNCTION: PROTEIN C IS A VITAMIN K-DEPENDENT SERINE PROTEASE THAT
CC REGULATES BLOOD COAGULATION BY INACTIVATING FACTORS VA AND VIIIA
CC IN THE PRESENCE OF CALCIUM IONS AND PHOSPHOLIPIDS.
CC -1- CATALYTIC ACTIVITY: Degradation of blood coagulation factors Va
CC and VIIIA.
CC -1- SUBUNIT: SYNTHESIZED AS A SINGLE CHAIN PRECURSOR, WHICH IS CLEAVED
CC INTO A LIGHT CHAIN AND A HEAVY CHAIN HELD TOGETHER BY A DISULFIDE
CC BOND. THE ENZYME IS THEN ACTIVATED BY THROMBIN, WHICH CLEAVES A
CC TETRADECAPEPTIDE FROM THE AMINO END OF THE HEAVY CHAIN; THIS
CC REACTION, WHICH OCCURS AT THE SURFACE OF ENDOTHELIAL CELLS, IS
CC STRONGLY PROMOTED BY THROMBOMODULIN.
CC -1- TISSUE SPECIFICITY: PLASMA; SYNTHESIZED IN THE LIVER.
CC -1- PTM: THE VITAMIN K-DEPENDENT ENZYMIC CARBOXYLATION OF SOME
CC GLU RESIDUES ALLOWS THE MODIFIED PROTEIN TO BIND CALCIUM.
CC -1- MISCELLANEOUS: CALCIUM ALSO BINDS, WITH STRONGER AFFINITY TO
CC ANOTHER SITE, BEYOND THE GLA DOMAIN. THIS GLA-INDEPENDENT BINDING
CC SITE IS NECESSARY FOR THE RECOGNITION OF THE
CC THROMBIN-THROMBOMODULIN COMPLEX.
CC -1- SIMILARITY: BELONGS TO PEPTIDASE FAMILY S1.
CC -1- SIMILARITY: CONTAINS 2 EGF-LIKE DOMAINS.
CC -----
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CC -----
DR EMBL: U49933; AAA92956.1; -
DR HSSP: P04070; 1PCU.
DR MEROPS: S01.218; -
DR InterPro: IPR000152; Asx_hydroxyl.
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DR InterPro: IPR000294; Vitk_dep_GLA.
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DR PROSITE: PS01187; EGF_Ca; 1.
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DR PROSITE: PS00135; TRYPSIN_SER; 1.
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KW Gamma-carboxyglutamic acid; Calcium-binding; Vitamin K; Hydroxylation;
KW EGF-like domain; Repeat; Endothelial cell; Hydrolase; Signal.
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Query Match 70.1%; Score 138; DB 1; Length 458;
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Matches 26; Conservative 4; Mismatches 14; Indels 0; Gaps 0;

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	DR	PROSITE; PS00022; EGF-1; 1.
	DR	PROSITE; PS01186; EGF-2; 2.
	DR	PROSITE; PS01187; EGF_CA_1.
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	DR	PROSITE; PS00240; TRYPSIN_DOM; 1.
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KW	Blood coagulation; Glycoprotein; Serine protease;	
KM	Gamma-carboxyglutamic acid; Calcium-binding; Vitamin K Hydroxylation;	
KV	EGF-like domain; Repeat; Endothelial cell; Hydrolase; Signal.	
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FT	PROPEP	19 41
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FT	DISULFD	100 105
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FT	DISULFD	161 174
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FT	DISULFD	396 424
FT	CARBOHYD	138 138
FT	CARBOHYD	292 292
FT	CARBOHYD	353 353
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Query Match	Best Local Similarity	62.4%; Score 123; DB 1; Length 459;
Matches	23; Conservative	Pred. No. 2.2e-13; 7; Mismatches 14; Indels 0; Gaps 0;





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FT DISULFID 159 172 BY SIMILARITY.
FT DISULFID 180 318 INTERCHAIN.
FT DISULFID 237 253
FT DISULFID 368 382
FT DISULFID 393 421
FT CARBOHYD 136 136 N-LINKED (GLCNAC. . .)
FT CARBOHYD 289 289 N-LINKED (GLCNAC. . .)
FT CARBOHYD 350 350 N-LINKED (GLCNAC. . .)
FT CARBOHYD 366 366 N-LINKED (GLCNAC. . .)
FT VARIANT 82 82 F->K.
FT CONFLICT 455 456 VP->PV (IN REF. 4).
SQ SEQUENCE 456 AA: 51407 MM: CAA6833F894C209 CRC64;

Query Match 61.9% Score 122: DB 1: Length 456;
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Matches 21: Conservative 9; Mismatches 12; Indels 0; Gaps 0;

OY 1 ANSFLXLRQSLRXKXCIXXICDFXAKXIFEDVDPTLAWS 42
Db 40 ANSFLLELRGNYRRCSEVCEFEAREIFONTEDTMFWS 81

RESULT 7
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ID FA10_BOVIN
AC P00743:
DT 21-JUL-1986 (Rel. 01, Created)
DT 13-AUG-1987 (Rel. 05, Last sequence update)
DT 15-JUN-2002 (Rel. 41, Last annotation update)
DE Coagulation factor X precursor (EC 3.4.21.6) (Stuart factor).
GN F10.
OS Bos taurus (Bovine).
OC Eukaryota; Metazoa; Chordata; Vertebrata; Euteleostomi;
OC Mammalia; Eutheria; Cetartiodactyla; Ruminantia; Pecora; Bovidae;
OC Bovidae; Bovinae; Bos.
OX NCBI_TaxID=9913;
RN [1]
RP MEDLINE=84247315; PubMed=6330671;
RX Fung M.R., Campbell R.M., McGilivray R.T.A.;
RT "Blood coagulation factor X mRNA encodes a single polypeptide chain
RL containing a prepro leader sequence."
RL Nucleic Acids Res. 12:4481-4492(1984).
RN [2]
RP SEQUENCE OF 41-180.
RX MEDLINE=80130563; PubMed=6766735;
RA Enfield D.L., Ericsson L.H., Fujikawa K., Walsh K.A., Neurath H.,
RA Titani K.;
RT "Amino acid sequence of the light chain of bovine factor XI (Stuart
RT factor)."
RL Biochemistry 19:659-667(1980).
RN [3]
RP REVISION TO 103.
RX MEDLINE=83308813; PubMed=668526;
RA McMillen B.A., Fujikawa K., Kistel W.;
RT "The occurrence of beta-hydroxyaspartic acid in the vitamin
RT K-dependent blood coagulation zymogens."
RL Biochem. Biophys. Res. Commun. 115:8-14(1993).
RN [4]
RP SEQUENCE OF 183-492, CARBOHYDRATE-LINKAGE SITES, AND DISULFIDE BONDS.
RX MEDLINE=76033069; PubMed=1059093;
RA Titani K., Fujikawa K., Enfield D.L., Ericsson L.H., Walsh K.A.,
RA Neurath H.;
RT "Bovine factor XI (Stuart factor): amino-acid sequence of heavy
RT chain."
RL Proc. Natl. Acad. Sci. U.S.A. 72:3082-3086(1975).
RN [5]
RP SEQUENCE OF 183-233, AND CARBOHYDRATE-LINKAGE SITES.
RX MEDLINE=94062825; PubMed=8243461;
RA Inoue K., Morita T.;
RT "Identification of O-linked oligosaccharide chains in the activation
RT peptides of blood coagulation factor X. The role of the carbohydrate
RT moieties in the activation of factor X."

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RL Eur. J. Biochem. 218:153-163(1993).
RN [6]
RP ACTIVE SITE.
RX MEDLINE=73053314; PubMed=4264286;
RA Titani K., Hermanson M.A., Fujikawa K., Ericsson L.H., Walsh K.A.,
RA Neurath H., Davie E.W.;
RT "Bovine factor XIa (activated Stuart factor). Evidence of homology
RT with mammalian serine proteases."
RL Biochemistry 11:4899-4903(1972).
RN [7]
RP PROCESSING.
RX MEDLINE=76053121; PubMed=1059122;
RA Fujikawa K., Titani K., Davie E.W.;
RT "Activation of bovine factor X (Stuart factor): conversion of factor
RT Xa-alpha to factor Xa-beta."
RL Proc. Natl. Acad. Sci. U.S.A. 72:3359-3363(1975).
RN [8]
RP CALCIUM-BINDING DATA.
RX MEDLINE=84185716; PubMed=6546930;
RA Sugo T., Björk I., Holmgren A., Stenflo J.;
RT "Calcium-binding properties of bovine factor X lacking the gamma-
RT carboxyglutamic acid-containing region."
RL J. Biol. Chem. 259:5705-5710(1984).
RN [9]
RP SULFATON.
RX MEDLINE=86140210; PubMed=3949800;
RA Morita T., Jackson C.M.;
RT "Localization of the structural difference between bovine blood
RT coagulation factors XI and X2 to tyrosine 18 in the activation
RT peptide."
RL J. Biol. Chem. 261:4008-4014(1986).
RN [10]
RP STRUCTURE BY NMR OF 85-126.
RX MEDLINE=91084483; PubMed=2261466;
RA Selander M., Persson E., Stenflo J., Drakenberg T.;
RT "1H NMR assignment and secondary structure of the Ca2(+)-free form of
RT the amino-terminal epidermal growth factor like domain in coagulation
RL factor X."
RL Biochemistry 29:8111-8118(1990).
RN [11]
RP STRUCTURE BY NMR OF 85-126.
RX MEDLINE=92329412; PubMed=1627540;
RA Ullner M., Selander M., Persson E., Stenflo J., Drakenberg T.,
RA Telemann O.;
RT "Three-dimensional structure of the apo form of the N-terminal
RT EGF-like module of blood coagulation factor X as determined by NMR
RL spectroscopy and simulated folding."
RL Biochemistry 31:5974-5983(1992).
RN [12]
RP STRUCTURE BY NMR OF 85-126.
RX MEDLINE=92406922; PubMed=1527084;
RA Selander M., Ullner M., Persson E., Telemann O.,
RA Stenflo J., Drakenberg T.;
RT "How an epidermal growth factor (EGF)-like domain binds calcium. High
RT resolution NMR structure of the calcium form of the NH2-terminal EGF-
RT like domain in coagulation factor X."
RL J. Biol. Chem. 267:19642-19649(1992).
RN [13]
RP STRUCTURE BY NMR OF 41-126.
RX MEDLINE=96387194; PubMed=8794734;
RA Sunnerhagen M., Olah G.A., Stenflo J., Forsen S., Drakenberg T.,
RA Trewheella J.;
RT "The relative orientation of Glu and EGF domains in coagulation
RT factor X is altered by Ca2+ binding to the first EGF domain. A
RT combined NMR-small angle x-ray scattering study."
RL Biochemistry 35:11547-11559(1996).
RN [14]
RP FUNCTION: Factor Xa is a vitamin K-dependent glycoprotein that
RP converts prothrombin to thrombin in the presence of factor Va,
RP calcium and phospholipid during blood clotting.
CC -1- CATALYTIC ACTIVITY: Preferential cleavage: Arg-|-Thr and then
CC Arg-|-Ile bonds in prothrombin to form thrombin.
CC -1- SUBUNIT: THE TWO CHAINS ARE FORMED FROM A SINGLE-CHAIN PRECURSOR
CC BY THE EXCISION OF TWO ARG RESIDUES AND ARE HELD TOGETHER BY 1 OR

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CC MORE DISULFIDE BONDS.  
 CC -1- PTM: THE VITAMIN K-DEPENDENT, ENZYMATIC CARBOXYLATION OF SCHE  
 CC GLUTAMIC ACID RESIDUES ALLOWS THE MODIFIED PROTEIN TO BIND  
 CC CALCIUM.  
 CC -1- PTM: N- AND O-GLYCOSYLATED.  
 CC -1- PTM: THE ACTIVATION PEPTIDE IS CLEAVED BY FACTOR IXA (IN THE  
 CC INTRINSIC PATHWAY), OR BY FACTOR VIIA (IN THE EXTRINSIC PATHWAY).  
 CC MISCELLANEOUS: CALCIUM ALSO BINDS, WITH STRONGER AFFINITY TO  
 CC ANOTHER SITE, BEYOND THE GLA DOMAIN.  
 CC -1- SIMILARITY: BELONGS TO PEPTIDASE FAMILY S1.  
 CC -1- SIMILARITY: CONTAINS 2 EGF-LIKE DOMAINS.  
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 CC EMBL: X00673; CAA25286.1; -  
 CC PIR: A00925; EXBO. -  
 CC PDB: 1APO; 31-JAN-94.  
 CC PDB: 1CCF; 31-MAY-94.  
 CC PDB: 1MHE; 15-MAY-97.  
 CC PDB: 1MHE; 15-MAY-97.  
 CC MEROPS: S01.216; -  
 CC GlycosultedB: P00743; -  
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 CC InterPro: IPR001314; Chymotrypsin.  
 CC InterPro: IPR000561; EGF-1-like.  
 CC InterPro: IPR000742; EGF-2.  
 CC InterPro: IPR001881; EGF\_Ca.  
 CC InterPro: IPR002383; GLA\_blood.  
 CC InterPro: IPR001254; Ser\_protease\_Try.  
 CC InterPro: IPR000294; Vitr\_dep\_GLA.  
 CC Pfam: PF00008; EGF\_2.  
 CC Pfam: PF00089; trypsin\_1.  
 CC Pfam: PF00594; gla\_1.  
 CC PRINTS: PR00722; CHYMOTRYPSIN.  
 CC PRINTS: PR00001; GLABLOOD.  
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 CC SMART: SM00001; EGF\_1like; 1.  
 CC SMART: SM00069; GLA; 1.  
 CC SMART: SM00020; Tryp\_SPC; 1.  
 CC PROSITE: PS00010; ASX\_HYDROXYL; 1.  
 CC PROSITE: PS00022; EGF\_1; 1.  
 CC PROSITE: PS01186; EGF\_2; 2.  
 CC PROSITE: PS01187; EGF\_CA; 1.  
 CC PROSITE: PS00011; GLU CARBOXYLATION; 1.  
 CC PROSITE: PS0240; TRYPSIN\_DOM; 1.  
 CC PROSITE: PS00134; TRYPSIN\_HIS; 1.  
 CC PROSITE: PS00135; TRYPSIN\_SER; 1.  
 CC Glycoprotein: Hydroxylase; Serine protease; Plasma; blood coagulation;  
 CC Gamma-carboxyglutamic acid; Hydroxylation; Calcium-binding; Vitamin K;  
 CC Signal; Zymogen; EGF-like domain; Repeat; Sulfation; 3D-structure.  
 FT SIGNAL 1 23  
 FT PROPEP 24 40  
 FT CHAIN 41 180  
 FT CHAIN 183 492  
 FT PROPEP 183 233  
 FT CHAIN 234 492  
 FT PROPEP 476 492  
 FT DOMAIN 86 122  
 FT DOMAIN 125 165  
 FT DOMAIN 234 492  
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 FT ACT\_SITE 418 418  
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 FT MOD\_RES 47 47  
 FT MOD\_RES 54 54  
 FT MOD\_RES 54 54

FT MOD\_RES 56 56 GAMMA-CARBOXYGLUTAMIC ACID.  
 FT MOD\_RES 59 59 GAMMA-CARBOXYGLUTAMIC ACID.  
 FT MOD\_RES 60 60 GAMMA-CARBOXYGLUTAMIC ACID.  
 FT MOD\_RES 65 65 GAMMA-CARBOXYGLUTAMIC ACID.  
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 QY 1 ANSFLLXLRGSGLRXCIXXICFXXAKXIFEDVDTLAWSKH 44  
 DB 41 ANSFLEVKQGNLDRCLERACSLERAREVEDEAEDTDEPMKY 84  
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 ID FA10\_HUMAN STANDARD; PRT; 488 AA.  
 AC P00742; Q14340;  
 DT 21-JUL-1986 (Rel. 01, Created)  
 DT 01-OCT-1989 (Rel. 12, Last sequence update)  
 DT 15-JUN-2002 (Rel. 41, Last annotation update)  
 DE Coagulation factor X precursor (EC 3.4.21.6) (Stuart factor).  
 GN F10.  
 OS Homo sapiens (Human).  
 OC Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
 OC Mammalia; Eutheria; Primates; Catarrhini; Homnidae; Homo.  
 OX NCBI\_TaxID=9606;  
 RN [1]  
 RN SEQUENCE FROM N.A.  
 RX MEDLINE=91216473; PubMed=1902434;  
 RA Messier T.L., Pittman D.D., Long G.L., Kaufman R.J., Church W.R.;  
 RT "Cloning and expression in COS-1 cells of a full-length cDNA encoding  
 RL human coagulation factor X.";  
 RL Gene 99:291-294 (1991).  
 RN [2]  
 RN SEQUENCE FROM N.A.  
 RX MEDLINE=87026600; PubMed=3768336;  
 RA Leytus S.P., Foster D.C., Kurachi K., Davie E.W.;  
 RT "Gene for human factor X: a blood coagulation factor whose gene  
 RL organization is essentially identical with that of factor IX and  
 RL protein C.";  
 RL Biochemistry 25:5098-5102 (1986).  
 RN [3]  
 RN SEQUENCE OF 13-488 FROM N.A.  
 RX MEDLINE=85216545; PubMed=2582420;  
 RA Fung W.R., Hay C.W., McGillivray R.T.A.;  
 RT "Characterization of an almost full-length cDNA coding for human  
 RL blood coagulation factor X.";  
 RL Proc. Natl. Acad. Sci. U.S.A. 82:3591-3595 (1985).  
 RN [4]  
 RN SEQUENCE OF 19-488 FROM N.A.  
 RC TISSUE=Liver;  
 RX MEDLINE=86221713; PubMed=3011603;  
 RA Kaul R.K., Hildebrand B., Roberts S., Jagadeeswaran P.;  
 RT "Isolation and characterization of human blood-coagulation factor X  
 RL cDNA.";  
 RL Gene 41:311-314 (1986).  
 RN [5]  
 RN SEQUENCE OF 41-179.  
 RX MEDLINE=83257207; PubMed=6871167;  
 RA McMullen B.A., Fujikawa K., Kisiel W., Sasagawa T., Howald W.N.,  
 RA Kwa E.Y., Weinstein B.;  
 RT "Complete amino acid sequence of the light chain of human blood  
 RL coagulation factor X: evidence for identification of residue 63 as  
 RL beta-hydroxyaspartic acid.";  
 RL Biochemistry 22:2875-2884 (1983).  
 RN [6]  
 RN SEQUENCE OF 115-488 FROM N.A., AND TISSUE SPECIFICITY.  
 RC TISSUE=Liver;  
 RX MEDLINE=84222026; PubMed=6587384;  
 RA Leytus S.P., Chung D.W., Kisiel W., Kurachi K., Davie E.W.;  
 RT "Characterization of a cDNA coding for human factor X.";

Proc. Natl. Acad. Sci. U.S.A. 81:3699-3702(1984).  
 (17)  
 RN SEQUENCE OF 183-234, AND CARBOHYDRATE-LINKAGE SITES.  
 RX MEDLINE-94062825; PubMed-8243461;  
 RA Inoue K., Morita T.,  
 RT "Identification of O-linked oligosaccharide chains in the activation  
 RT peptides of blood coagulation factor X. The role of the carbohydrate  
 RT moieties in the activation of factor X.";  
 RL Eur. J. Biochem. 218:153-163(1993).  
 (18)  
 RN SEQUENCE OF 1-23 FROM N.A.  
 RX MEDLINE-90128299; PubMed-2612918;  
 RA Jagadeeswaran P., Reddy S.V., Rao K.J., Hamsabhusanam K., Lyman G.,  
 RT "Cloning and characterization of the 5' end (exon 1) of the gene  
 RT encoding human factor X.";  
 RL Gene 84:517-519(1989).  
 (19)  
 RN X-RAY CRYSTALLOGRAPHY (2.2 ANGSTROMS) OF 86-179 AND 235-278.  
 RX MEDLINE-93360277; PubMed-8355279;  
 RA Padmanabhan K., Padmanabhan K.P., Tulinsky A., Park C.H., Bode W.,  
 RT Huber R., Blankenship D.T., Cardin A.D., Kistiel W.,  
 RT "Structure of human des(1-45) factor Xa at 2.2-A resolution.";  
 RL J. Mol. Biol. 232:947-966(1993).  
 (10)  
 RN X-RAY CRYSTALLOGRAPHY (2.3 ANGSTROMS) OF 86-179 AND 235-278.  
 RX MEDLINE-98283982; PubMed-9618463;  
 RA Kanata K., Kawamoto H., Honma T., Iwama T., Kim S.H.,  
 RT "Structural basis for chemical inhibition of human blood coagulation  
 RT factor Xa.";  
 RL Proc. Natl. Acad. Sci. U.S.A. 95:6630-6635(1998).  
 (1)  
 CC -1- FUNCTION: Factor Xa is a vitamin K-dependent glycoprotein that  
 CC converts prothrombin to thrombin in the presence of factor Va,  
 CC calcium and phospholipid during blood clotting.  
 CC -1- CATALYTIC ACTIVITY: Preferential cleavage: Arg-|-Thr and then  
 CC Arg-|-Ile bonds in prothrombin to form thrombin.  
 CC -1- SUBUNIT: THE TWO CHAINS ARE FORMED FROM A SINGLE-CHAIN PRECURSOR  
 CC BY THE EXCISION OF TWO ARG RESIDUES AND ARE HELD TOGETHER BY 1 OR  
 CC MORE DISULFIDE BONDS.  
 CC -1- TISSUE SPECIFICITY: Plasma; synthesized in the liver.  
 CC -1- PM: THE VITAMIN K-DEPENDENT, ENZYMAIC CARBOXYLATION OF SOME  
 CC GLUTAMIC ACID RESIDUES ALLOWS THE MODIFIED PROTEIN TO BIND  
 CC CALCIUM.  
 CC -1- PM: N- AND O-GLYCOSYLATED.  
 CC -1- INTRINSIC PATHWAY), OR BY FACTOR VIIA (IN THE EXTRINSIC PATHWAY).  
 CC -1- SIMILARITY: BELONGS TO PEPTIDASE FAMILY S1  
 CC -1- SIMILARITY: CONTAINS 2 EGF-LIKE DOMAINS.  
 CC -----  
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 CC -----  
 DR EMBL: K03194; AAA52490.1; -  
 DR EMBL: M57285; AAA52421.1; -  
 DR EMBL: L29433; AAA52764.1; -  
 DR EMBL: L00390; AAA52764.1; JOINED.  
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 DR EMBL: L00395; AAA52764.1; JOINED.  
 DR EMBL: L00396; AAA52764.1; JOINED.  
 DR EMBL: M22613; AAA51984.1; -  
 DR EMBL: K01886; AAA52486.1; -  
 DR EMBL: M33297; AAA52636.1; -  
 DR PIR: A00924; EXHU.  
 DR PIR: A25853; A25853.  
 DR PIR: A24478; A24478.  
 DR PDB: IHCG; 08-MAY-95.

DR PDB: 1FXA; 29-OCT-97.  
 DR PDB: 1EXY; 17-OUN-98.  
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 DR Gene: HGNC:3528; F10.  
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 DR InterPro: IPR000561; EGF-like.  
 DR InterPro: IPR000742; EGF-2.  
 DR InterPro: IPR001881; EGF-Ca.  
 DR InterPro: IPR002383; GLA\_blood.  
 DR InterPro: IPR001254; Ser\_protease\_Try.  
 DR InterPro: IPR000294; Vitk\_dep\_GLA.  
 DR Pfam: PF00089; EGF; 2.  
 DR Pfam: PF00089; trypsin; 1.  
 DR Pfam: PF00594; gla; 1.  
 DR PRINTS: PR00722; CHYMOTRYPSIN.  
 DR PRINTS: PR00001; GLABLOOD.  
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 DR PROSITE: PS01187; EGF\_CA; 1.  
 DR PROSITE: PS00011; GLU\_CARBOXYLATION; 1.  
 DR PROSITE: PS02040; TRYPSIN\_DOM; 1.  
 DR PROSITE: PS00134; TRYPSIN\_HIS; 1.  
 DR PROSITE: PS00135; TRYPSIN\_SER; 1.  
 DR GlycoSuiteDB: Serine protease; Plasma; Blood coagulation;  
 DR Gamma-carboxyglutamic acid; Hydroxylation; Calcium-binding; Vitamin K;  
 DR Signal; zymogen; EGF-like domain; Repeat; 3D-structure.  
 DR Signal; 1  
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 FT DISULFID 112 121  
 FT DISULFID 129 140  
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KW Gamma-carboxyglutamic acid; Hydroxylation; Calcium-binding; Vitamin K;  
 KW Signal; Zymogen; EGF-like domain; Repeat.  
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 FT CHAIN 86 122  
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 FT DISULFID 413 441  
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 FT CARBOHYD 187 187  
 FT CARBOHYD 205 205  
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Query Match 52.3%; Score 103; DB 1; Length 490;  
 Best Local Similarity 43.2%; Pred. No. 6; Be-10;  
 Matches 19; Conservative 9; Mismatches 16; Indels 0; Gaps 0;

OY 1 ANSFLXLRGSLKXKICIXICDXXAKXIPEDVDLTAFWSKH 44  
 Db 41 ANSFLXLRGSLKXKICIXICDXXAKXIPEDVDLTAFWSKH 44

RESULT 11  
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 AC P98139; P79224;  
 DT 01-FEB-1996 (Rel. 33, Created)  
 DT 15-JUN-1998 (Rel. 36, Last sequence update)  
 DT 15-JUN-2002 (Rel. 41, Last annotation update)  
 DE Coagulation factor VII precursor (EC 3.4.21.21) (Serum prothrombin conversion accelerator).

GN F7.  
 OS *Oryctolagus cuniculus* (Rabbit).  
 OC Eukaryota; Metazoa; Chordata; Craniata; Euteleostomi;  
 OC Mammalia; Eutheria; Lagomorpha; Leporidae; Oryctolagus.  
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 RP [1]  
 RP SEQUENCE FROM N.A.  
 RC TISSUE=Liver;  
 RA MEDLINE=93190306; PubMed=8383365;  
 RX Brochers A.B., Clarke B.J., Sheffield W.P., Blajchman M.A.;  
 RT "Complete nucleotide sequence of the cDNA encoding rabbit coagulation factor VII";  
 RL Thromb. Res. Suppl. 69:231-238(1993).  
 RN [2]  
 RP REVISION TO 395.  
 RC TISSUE=Liver;  
 RA Ruiz S.R., Blajchman M.A., Clarke B.J.;  
 RL Submitted (NOV-1996) to the EMBL/Genbank/DBJ databases.  
 CC -1- FUNCTION: CIRCULATES IN THE BLOOD IN A ZYMOGEN FORM. FACTOR VII IS CONVERTED TO FACTOR VIIA BY FACTOR XA, FACTOR XIIA, FACTOR IXA, OR THROMBIN BY MINOR PROTEOLYSIS. IN THE PRESENCE OF TISSUE FACTOR AND CALCIUM IONS, FACTOR VIIA THEN CONVERTS FACTOR X TO FACTOR XA BY LIMITED PROTEOLYSIS. FACTOR VIIA WILL ALSO CONVERT FACTOR IX TO FACTOR IXA IN THE PRESENCE OF TISSUE FACTOR AND CALCIUM (BY SIMILARITY).  
 CC -1- CATALYTIC ACTIVITY: Hydrolyzes one Arg-1-Ile bond in factor X to form factor Xa.  
 CC -1- SUBUNIT: HETERODIMER OF A LIGHT CHAIN AND A HEAVY CHAIN LINKED BY A DISULFIDE BOND (BY SIMILARITY).  
 CC -1- TISSUE SPECIFICITY: PLASMA.  
 CC -1- PM: THE VITAMIN K-DEPENDENT, ENZYMIC CARBOXYLATION OF SOME GLUTAMIC ACID RESIDUES ALLOWS THE MODIFIED PROTEIN TO BIND CALCIUM (BY SIMILARITY).  
 CC -1- SIMILARITY: BELONGS TO PEPTIDASE FAMILY S1.  
 CC -1- SIMILARITY: CONTAINS 2 EGF-LIKE DOMAINS.  
 CC -----  
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 CC -----  
 CC EMBL: U77477; AAB37326.1; -  
 DR HSSP; P08709; IFAK.  
 DR MEROPS; S01.215; -  
 DR InterPro: IPR000152; Asx\_hydroxyl.  
 DR InterPro: IPR001314; Chymotrypsin.  
 DR InterPro: IPR000561; EGF-like.  
 DR InterPro: IPR000742; EGF\_2.  
 DR InterPro: IPR001881; EGF\_Ca.  
 DR InterPro: IPR002383; GLA\_blood.  
 DR InterPro: IPR001254; Ser.protease\_Try.  
 DR InterPro: IPR000294; Vitk\_dep\_GLA.  
 DR Pfam; PF00008; EGF\_2.  
 DR Pfam; PF00089; Trypsin\_1.  
 DR Pfam; PF00594; gla\_1.  
 DR PRINTS; PR00722; CHYMOTRYPSIN.  
 DR PRINTS; PR00001; GLABLOOD.  
 DR SMART; SM00179; EGF\_CA\_1.  
 DR SMART; SM00001; EGF\_Like\_1.  
 DR SMART; SM00069; GLA\_1.  
 DR SMART; SM00020; TYP\_SPE\_1.  
 DR PROSITE; PS00010; ASX\_HYDROXYL\_1.  
 DR PROSITE; PS00022; EGF\_1\_1.  
 DR PROSITE; PS01186; EGF\_2\_1.  
 DR PROSITE; PS01187; EGF\_CA\_1.  
 DR PROSITE; PS00011; GLU\_CARBOXYLATION\_1.  
 DR PROSITE; PS50240; TRYPSIN\_DOM\_1.  
 DR PROSITE; PS00134; TRYPSIN\_HIS\_1.  
 DR PROSITE; PS00135; TRYPSIN\_SER\_1.  
 DR Hydrolase; Serine protease; Blood coagulation; Zymogen; Glycoprotein;

KM Liver; Plasma; Vitamin K; Calcium-binding; Gamma-carboxyglutamic acid;  
 KM EGF-like domain; Repeat; Signal; Hydroxylation.  
 FT SIGNAL 1 21 POTENTIAL.  
 FT PROPEP 22 39 POTENTIAL.  
 FT CHAIN 40 191 FACTOR VII LIGHT CHAIN.  
 FT CHAIN 192 444 FACTOR VII HEAVY CHAIN.  
 FT DOMAIN 45 74 GLA-RICH.  
 FT DOMAIN 85 121 EGF-LIKE 1, CALCIUM-BINDING (POTENTIAL).  
 FT DOMAIN 126 167 EGF-LIKE 2.  
 FT DOMAIN 192 444 SERINE PROTEASE.  
 FT SITE 191 192 CLEAVAGE (BY FACTOR Xa, FACTOR XIIa, FACTOR IXa, OR THROMBIN) (BY SIMILARITY).  
 FT ACT\_SITE 232 232 BY SIMILARITY.  
 FT ACT\_SITE 281 281 BY SIMILARITY.  
 FT ACT\_SITE 383 383 BY SIMILARITY.  
 FT BINDING 377 377 SUBSTRATE (BY SIMILARITY).  
 FT BINDING 377 377 BY SIMILARITY.  
 FT DISULFID 56 61 BY SIMILARITY.  
 FT DISULFID 89 100 BY SIMILARITY.  
 FT DISULFID 94 109 BY SIMILARITY.  
 FT DISULFID 111 120 BY SIMILARITY.  
 FT DISULFID 130 141 BY SIMILARITY.  
 FT DISULFID 137 151 BY SIMILARITY.  
 FT DISULFID 153 166 BY SIMILARITY.  
 FT DISULFID 174 301 BY SIMILARITY.  
 FT DISULFID 198 203 BY SIMILARITY.  
 FT DISULFID 217 233 BY SIMILARITY.  
 FT DISULFID 349 368 BY SIMILARITY.  
 FT DISULFID 379 407 BY SIMILARITY.  
 FT MOD\_RES 45 45 GAMMA-CARBOXYGLUTAMIC ACID.  
 FT MOD\_RES 46 46 GAMMA-CARBOXYGLUTAMIC ACID.  
 FT MOD\_RES 53 53 GAMMA-CARBOXYGLUTAMIC ACID.  
 FT MOD\_RES 55 55 GAMMA-CARBOXYGLUTAMIC ACID.  
 FT MOD\_RES 58 58 GAMMA-CARBOXYGLUTAMIC ACID.  
 FT MOD\_RES 59 59 GAMMA-CARBOXYGLUTAMIC ACID.  
 FT MOD\_RES 64 64 GAMMA-CARBOXYGLUTAMIC ACID.  
 FT MOD\_RES 65 65 GAMMA-CARBOXYGLUTAMIC ACID.  
 FT MOD\_RES 68 68 GAMMA-CARBOXYGLUTAMIC ACID.  
 FT MOD\_RES 74 74 GAMMA-CARBOXYGLUTAMIC ACID.  
 FT MOD\_RES 102 102 HYDROXYLATION (BY SIMILARITY).  
 FT MOD\_RES 211 211 N-LINKED (GLCNAC. . .) (POTENTIAL).  
 FT CARBOHYD 211 211 N-LINKED (GLCNAC. . .) (POTENTIAL).  
 FT CARBOHYD 306 306 N-LINKED (GLCNAC. . .) (POTENTIAL).  
 SQ SEQUENCE 444 AA; 49011 MW; 0481ABC4FE5427F8 CRC64;  
 Query Match 51.3%; Score 101; DB 1; Length 444;  
 Best Local Similarity 46.3%; Pred. NO. 1.3e-09;  
 Matches 19; Conservative 5; Mismatches 17; Indels 0; Gaps 0;

Db 1 ANSFLXLRGSLRXICIXXICDPXKXKXIFEDVDLAFW 41  
 40 ANSFLLELRPGSLERCKEKLCSFEARAREVFQSTERTKQFW 80

RESULT 12  
 FAY\_HUMAN STANDARD; PRT; 466 AA.  
 ID FAY\_HUMAN 014339;  
 AC P08709; 014339;  
 DT 01-JAN-1988 (Rel. 06, Created)  
 DT 01-JAN-1988 (Rel. 06, Last sequence update)  
 DT 15-JUN-2002 (Rel. 41, Last annotation update)  
 DE Coagulation factor VII precursor (EC 3.4.21.21) (Serum prothrombin conversion accelerator) (Eptacog alfa).  
 DE CN  
 OS Homo sapiens (Human).  
 OC Eukaryota; Metazoa; Chordata; Vertebrata; Euteleostomi;  
 OC Mammalia; Eutheria; Primates; Catarrhini; Homiidae; Homo.  
 OX NCBI\_TaxID:9606;  
 RN [1]  
 RC SEQUENCE FROM N.A.  
 RC TISSUE=Liver;  
 RX MEDLINE-86205965; PubMed-3486420;  
 RA Hagen F.S., Gray C.L., O'Hara P.J., Grant F.J., Saari G.C., Woodbury R.G., Hart C.E., Insley M.Y., Kistiel W., Kurachi K.,

RA Davie E.W.;  
 RT "Characterization of a cDNA coding for human factor VII.";  
 RT Proc. Natl. Acad. Sci. U.S.A. 83:2412-2416(1986).  
 RN [2]  
 RP SEQUENCE FROM N.A.  
 RX MEDLINE-87260948; PubMed-3037537;  
 RA O'Hara P.J., Grant F.J., Haldeman B.A., Gray C.L., Insley M.Y., Hagen F.S., Murray M.J.;  
 RT "Nucleotide sequence of the gene coding for human factor VII, a vitamin K-dependent protein participating in blood coagulation.";  
 RT Proc. Natl. Acad. Sci. U.S.A. 84:5158-5162(1987).  
 RN [3]  
 RP SEQUENCE FROM N.A., AND VARIANTS THR-352; GLN-413 AND LYS-445.  
 RA Rieger M.J., Armet T.Z., Carrington D.P., Chung M.-W., Lee K.L., Poel C.L., Toth E.J., Yi O., Nickerson D.A.;  
 RL Submitted (JAN-2002) to the EMBL/GenBank/DBJ databases.  
 RN [4]  
 RP SEQUENCE OF 61-466, AND POST-TRANSLATIONAL MODIFICATIONS.  
 RX MEDLINE-89088153; PubMed-3264725;  
 RA Thim L., Bjoern S., Christensen M., Nicolaisen E.M., Lund-Hansen T., Pedersen A.H., Hedner U.;  
 RT "Amino acid sequence and posttranslational modifications of human factor VIIa from plasma and transfected baby hamster kidney cells.";  
 RL Biochemistry 27:7785-7793(1988).  
 RN [5]  
 RP CARBOHYDRATE-LINKAGE SITES SER-112 AND SER-120.  
 RX MEDLINE-91250411; PubMed-1904059;  
 RA Bjoern S., Foster D.C., Thim L., Wiberg F.C., Christensen M., Komiyama Y., Pedersen A.H., Kistiel W.;  
 RT "Human plasma and recombinant factor VII. Characterization of O-glycosylations at serine residues 52 and 60 and effects of site-directed mutagenesis of serine 52 to alanine.";  
 RL J. Biol. Chem. 266:11051-11057(1991).  
 RN [6]  
 RP STRUCTURE OF CARBOHYDRATE ON SER-112.  
 RX MEDLINE-90062160; PubMed-2511201;  
 RA Nishimura H., Kawabata S., Kistiel W., Hase S., Ikenaka T., Takao T., Shimomishi Y., Iwanaga S.;  
 RT "Identification of a disaccharide (Xyl-Glc) and a trisaccharide (Xyl2-Glc) O-glycosidically linked to a serine residue in the first epidermal growth factor-like domain of human factors VII and IX and protein Z and bovine protein Z.";  
 RL J. Biol. Chem. 264:20320-20325(1989).  
 RN [7]  
 RP STRUCTURE OF CARBOHYDRATE ON SER-112.  
 RX MEDLINE-91344709; PubMed-2129367;  
 RA Iwanaga S., Nishimura H., Kawabata S., Kistiel W., Hase S., Ikenaka T.;  
 RT "A new trisaccharide sugar chain linked to a serine residue in the first EGF-like domain of clotting factors VII and IX and protein Z.";  
 RL Adv. Exp. Med. Biol. 281:121-131(1990).  
 RN [8]  
 RP X-RAY CRYSTALLOGRAPHY (2.0 ANGSTROMS) OF FVIIA IN COMPLEX WITH TF.  
 RX MEDLINE-96175641; PubMed-8598903;  
 RA Banner D.W., D'Arcy A., Chene C., Winkler F.K., Guha A., Kongsberg W.H., Nemerson Y., Kirchhofer D.;  
 RT "The crystal structure of the complex of blood coagulation factor VIIa with soluble tissue factor.";  
 RL Nature 380:41-46(1996).  
 RN [9]  
 RP X-RAY CRYSTALLOGRAPHY (2.1 ANGSTROMS) OF FVIIA IN COMPLEX WITH TF.  
 RX MEDLINE-99126538; PubMed-9925787;  
 RA Zhang E., St. Charles R., Tulinusky A.;  
 RT "Structure of extracellular tissue factor complexed with factor VIIa inhibited with a BPTI mutant.";  
 RL J. Mol. Biol. 285:2089-2104(1999).  
 RN [10]  
 RP STRUCTURE BY NMR OF 105-145.  
 RX MEDLINE-98367502; PubMed-9692950;  
 RA Muranyi A., Flinn B.E., Gippert G.P., Forsen S., Stenflo J., Drakenberg T.;  
 RT "Solution structure of the N-terminal EGF-like domain from human factor VII.";  
 RL Biochemistry 37:10605-10615(1998).



RL	Hum. Mutat.	8:108-115(1996).
RN	[20]	
RP	VARIANT VAL-304.	
RX	Tamary H., Fromovich Y., Shalmon L., Reich Z., Dym O., Lanir N.,	
RA	Brenner B., Paz M., Luder A.S., Blau O., Korostishevsky M.,	
RA	Zalizov R., Seligsohn U.;	
RT	"Ala244Val is a common, probably ancient mutation causing factor VII	
RT	deficiency in Moroccan and Iranian Jews.";	
RL	Thromb. Haemost.	76:283-291(1996).
RN	[21]	
RP	VARIANTS MALTA THR-194 AND VAL-304.	
RX	MEDLINE-98112461, PubMed-9452082;	
RA	Alshinawi C., Scerif C., Galdies R., Aquilina A., Felice A.E.;	
RT	"Two new missense mutations (P134F and A244V) in the coagulation	
RT	factor VII gene.";	
RL	Hum. Mutat. Suppl.	1:S189-S191(1998).
CC	-I- FUNCTIONED TO CIRCULATES IN THE BLOOD IN A ZYMOGEN FORM. FACTOR VII IS	
CC	CONVERTED TO FACTOR VIIA BY FACTOR XA, FACTOR XIIA, FACTOR IXA, OR	
CC	THROMBIN BY MINOR PROTEOLYSIS. IN THE PRESENCE OF TISSUE FACTOR	
CC	AND CALCIUM IONS, FACTOR VIIA THEN CONVERTS FACTOR X TO FACTOR XA	
CC	BY LIMITED PROTEOLYSIS. FACTOR VIIA WILL ALSO CONVERT FACTOR IX TO	
CC	FACTOR IXA IN THE PRESENCE OF TISSUE FACTOR AND CALCIUM.	
CC	-I- CATALYTIC ACTIVITY: Hydrolyzes one Arg-Ile bond in factor X to	
CC	form factor xa.	
CC	-I- SUBUNIT: HETEROIMER OF A LIGHT CHAIN AND A HEAVY CHAIN LINKED	
CC	BY A DISULFIDE BOND.	
CC	-I- ALTERNATIVE PRODUCTS: 2 isoforms; A (shown here) and B; are	
CC	produced by alternative splicing.	
CC	-I- TISSUE SPECIFICITY: PLASMA.	
CC	-I- PTM: THE VITAMIN K-DEPENDENT, ENZYMTIC CARBOXYLATION OF SOME	
CC	GLUTAMIC ACID RESIDUES ALLOWS THE MODIFIED PROTEIN TO BIND	
CC	CALCIUM.	
CC	-I- DISEASE: DEFECTS IN F7 CAN CAUSE COAGULOPATHY.	
CC	-I- PHARMACEUTICAL: Available under the names Nafasitase or Novoseven	
CC	(Novo Nordisk). Used for the treatment of bleeding episodes in	
CC		
Query Match	50.3%;	Score 99; DB 1; Length 466;
Best Local Similarity	48.8%;	Pred. No. 3.1e-09;
Matches 20; Conservative	4; Mismatches 17; Indels	0; Gaps 0;
Oy	1 ANSFPLXRQGSIXRXGTXICDFFKAXKXIFPDVDOTLAFW 41	
Db	61 ANAFLEELRPGSLERECKEBCGFEDAKRFDAERIKLFW 101	
	:   ::  :	
RESULT 13		
ID	TMGI_HUMAN	STANDARD: PRT; 218 AA.
AC	014668;	
DT	15-JUN-2002 (Rel. 41, Created)	
DT	15-JUN-2002 (Rel. 41, Last sequence update)	
DT	15-JUN-2002 (Rel. 41, Last annotation update)	
DE	Transmembrane gamma-carboxylutamic acid protein 1 precursor (Proline-	
DE	rich Gla protein 1) (Proline-rich gamma-carboxylutamic acid protein	
DE	1).	
GN	PRGCL OR TMGI OR PRGPL.	
OS	Homo sapiens (Human).	
OC	Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;	
OC	Mammalia; Eutheria; Primates; Catarrhini; Hominiidae; Homo.	
OX	NCBI_Taxid:9606;	
RN	[1]	
RP	SEQUENCE FROM N.A.	
RX	MEDLINE-97404347; PubMed-9256434.	
RA	Kulman J.D., Harris J.E., Haldeman B.A., Davie E.W.;	
RT	"Primary structure and tissue distribution of two novel proline-rich	
RT	gamma-carboxylutamic acid proteins.";	
CC	Proc. Natl. Acad. Sci. U.S.A. 94:9058-9062(1997).	
CC	-I- TISSUE SPECIFICITY: Highly expressed in the spinal cord.	
CC	-I- PTM: Gla residues are produced after subsequent posttranslational	
CC	modifications of glutamic acid by a vitamin K-dependent gamma-	
CC	carboxylase.	
CC		



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 CC -----  
 DR EMBL: AF009242; AAB67070.1; -  
 DR HSSP: P00740; ICRH.  
 DR Genew: HGNC:9469; PRRG1.  
 DR MIM: 604428; -  
 DR InterPro: IPR002383; GLA\_blood.  
 DR InterPro: IPR000294; VltK\_dep\_GLA.  
 DR Pfam: PF00594; gla; 1.  
 DR PRINTS: PR00001; GLABLOOD.  
 DR SMART: SM00069; GLA; 1.  
 DR PROSITE: PS00011; GLU-CARBOXYLATION; 1.  
 DR Gamma-carboxyglutamic acid; Vitamin K; Transmembrane.  
 FT PROPEP 1 20  
 FT CHAIN 21 218 TRANSMEMBRANE GAMMA-CARBOXYGLUTAMIC ACID  
 FT DOMAIN 21 83 EXTRACELLULAR (POTENTIAL).  
 FT TRANSMEM 84 106 POTENTIAL.  
 FT DOMAIN 107 218 CYTOPLASMIC (POTENTIAL).  
 FT DOMAIN 24 61 GLA-RICH.  
 FT DOMAIN 131 135 POLY-PRO.  
 FT SEQUENCE 218 AA: 24947 MW: 26538A61AB0AE98 CRC64;  
 SO  
 Query Match 46.7%; Score 92; DB 1; Length 218;  
 Best Local Similarity 38.6%; Pred. No. 2,1e-08;  
 Matches 17; Conservative 7; Mismatches 20; Indels 0; Gaps 0;  
 Oy 1 ANSFLXLRQGSIXRCIXICDFFXAKXIFEDVDITLAFMSKH 44  
 Db 21 ANGFEETIRQGNIERCKEECTFERAREAFENNEXTKEFWSTY 64  
 RESULT 14  
 ID THRB\_RAT STANDARD: PRT: 617 AA.  
 AC P18292;  
 DT 01-NOV-1990 (Rel. 16, Created)  
 DT 01-NOV-1990 (Rel. 16, Last sequence update)  
 DT 15-JUN-2002 (Rel. 41, Last annotation update)  
 DE Prothrombin precursor (EC 3.4.21.5).  
 GN CN  
 OS Rattus norvegicus (Rat).  
 OC Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
 OC Mammalia; Eutheria; Rodentia; Sciurognathi; Muridae; Murinae; Rattus.  
 OX NCBI\_Taxid=10116;  
 RN [1]  
 RP SEQUENCE FROM N.A.  
 RC STRAIN=Sprague-Dawley; TISSUE=Liver;  
 RA MEDLINE=90332426; PubMed=2377469;  
 RA Dihalich M., Monard D.;  
 RT "Nucleic Acids Res. 18:4251-4251 (1990).  
 RL [2]  
 RN SEQUENCE OF 363-617 FROM N.A.  
 RP TISSUE=Liver;  
 RC MEDLINE=92212913; PubMed=1557383;  
 RA Banfield D.K., Macgillivray R.T.;  
 RT "Partial characterization of vertebrate prothrombin cDNAs:  
 RT amplification and sequence analysis of the B chain of thrombin from  
 RT nine different species".  
 CC -1- FUNCTION: THROMBIN, WHICH CLEAVES BONDS AFTER ARG & LYS, CONVERTS  
 CC FIBRINOGEN TO FIBRIN AND ACTIVATES FACTORS V, VII, VIII, XIII,  
 CC AND, IN COMPLEX WITH THROMBOMODULIN, PROTEIN C.  
 CC -1- CATALYTIC ACTIVITY: Preferential cleavage: Arg-|-Gly; activates  
 CC fibrinogen to fibrin and releases fibrinopeptide A and B.  
 CC -1- PTM: THE GAMMA-CARBOXYGLUTAMYL RESIDUES, WHICH BIND CALCIUM IONS,

CC RESULT FROM THE CARBOXYLATION OF GLUTAMYL RESIDUES BY A MICROSOMAL  
 CC ENZYME. THE VITAMIN K-DEPENDENT CARBOXYLASE. THE MODIFIED RESIDUES  
 CC ARE NECESSARY FOR THE CA-DEPENDENT INTERACTION WITH A NEGATIVELY  
 CC CHARGED PHOSPHOLIPID SURFACE, WHICH IS ESSENTIAL FOR THE CONVERSION  
 CC OF PROTHROMBIN TO THROMBIN.  
 CC -1- MISCELLANEOUS: PROTHROMBIN IS ACTIVATED ON THE SURFACE OF A  
 CC PHOSPHOLIPID MEMBRANE THAT BINDS THE AMINO END OF PROTHROMBIN &  
 CC FACTORS VA & XA IN CA-DEPENDENT INTERACTIONS. FACTOR XA REMOVES  
 CC THE ACTIVATION PEPTIDE & CLEAVES THE REMAINING PART INTO LIGHT &  
 CC HEAVY CHAINS. THE ACTIVATION PROCESS STARTS SLOWLY BECAUSE FACTOR  
 CC V ITSELF HAS TO BE ACTIVATED BY THE INITIAL, SMALL AMOUNTS OF  
 CC THROMBIN.  
 CC -1- MISCELLANEOUS: THROMBIN CAN ITSELF CLEAVE THE AMINO TERMINAL  
 CC FRAGMENT (FRAGMENT 1) OF THE PROTHROMBIN, PRIOR TO ITS ACTIVATION  
 CC BY FACTOR XA.  
 CC -1- SIMILARITY: CONTAINS 2 KRINGLE DOMAINS.  
 CC -----  
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 CC -----  
 DR EMBL: X52835; CAA37017.1; -  
 DR EMBL: M81397; AAA42240.1; -  
 DR PIR: S10511; S10511.  
 DR HSSP: P00734; 1UWS.  
 DR MEROPS: S01.217; -  
 DR InterPro: IPR001314; Chymotrypsin.  
 DR InterPro: IPR002383; GLA\_blood.  
 DR InterPro: IPR000001; Kringle.  
 DR InterPro: IPR003966; Prothrombin.  
 DR InterPro: IPR001254; Ser.protease-try.  
 DR InterPro: IPR000294; VltK\_dep\_GLA.  
 DR Pfam: PF00051; kringle; 2.  
 DR Pfam: PF00089; trypsin; 1.  
 DR Pfam: PF00594; gla; 1.  
 DR PRINTS: PR00722; CHYMOTRYPSIN.  
 DR PRINTS: PR00001; GLABLOOD.  
 DR PRINTS: PR00018; KRINGLE.  
 DR PRINTS: PR01505; PROTHROMBIN.  
 DR ProDom: PD000395; Kringle; 2.  
 DR SMART: SM00069; GLA; 1.  
 DR SMART: SM00130; KR; 2.  
 DR SMART: SM00020; TRY-Spec; 1.  
 DR PROSITE: PS00011; GLU-CARBOXYLATION; 1.  
 DR PROSITE: PS00021; KRINGLE 1; 2.  
 DR PROSITE: PS00070; KRINGLE 2; 2.  
 DR PROSITE: PS00240; TRYPSIN\_DOM; 1.  
 DR PROSITE: PS00134; TRYPSIN\_HIS; 1.  
 DR PROSITE: PS00135; TRYPSIN\_SER; 1.  
 DR Blood coagulation; Plasma; Calcium-binding; Glycoprotein; Repeat;  
 KW Vitamin K; Zymogen; Gamma-carboxyglutamic acid; Acute phase; Liver;  
 KW Hydrolase; Serine protease; Kringle; Signal.  
 FT SIGNAL 1 24  
 FT PROPEP 25 43  
 FT CHAIN 44 617  
 FT PEPTIDE 44 200  
 FT PEPTIDE 201 323  
 FT CHAIN 324 359  
 FT CHAIN 360 617  
 FT DOMAIN 109 187  
 FT DOMAIN 215 292  
 FT DOMAIN 360 617  
 FT SITE 200 201  
 FT SITE 323 324  
 FT SITE 359 360  
 FT ACT\_SITE 402 402  
 FT ACT\_SITE 458 458  
 FT ACT\_SITE 564 564  
 PROTHROMBIN.  
 ACTIVATION PEPTIDE (FRAGMENT 1).  
 ACTIVATION PEPTIDE (FRAGMENT 2).  
 THROMBIN LIGHT CHAIN (A).  
 THROMBIN HEAVY CHAIN (B).  
 KRINGLE 1.  
 KRINGLE 2.  
 SERINE PROTEASE.  
 CLEAVAGE (BY THROMBIN).  
 CLEAVAGE (BY FACTOR XA).  
 CLEAVAGE (BY FACTOR XA).  
 CHARGE RELAY SYSTEM (BY SIMILARITY).  
 CHARGE RELAY SYSTEM (BY SIMILARITY).  
 CHARGE RELAY SYSTEM (BY SIMILARITY).

FT MOD\_RES 50 50 GAMMA-CARBOXYGLUTAMIC ACID.  
 FT MOD\_RES 51 51 GAMMA-CARBOXYGLUTAMIC ACID.  
 FT MOD\_RES 58 58 GAMMA-CARBOXYGLUTAMIC ACID.  
 FT MOD\_RES 60 60 GAMMA-CARBOXYGLUTAMIC ACID.  
 FT MOD\_RES 63 63 GAMMA-CARBOXYGLUTAMIC ACID.  
 FT MOD\_RES 64 64 GAMMA-CARBOXYGLUTAMIC ACID.  
 FT MOD\_RES 69 69 GAMMA-CARBOXYGLUTAMIC ACID.  
 FT MOD\_RES 70 70 GAMMA-CARBOXYGLUTAMIC ACID.  
 FT MOD\_RES 73 73 GAMMA-CARBOXYGLUTAMIC ACID.  
 FT MOD\_RES 76 76 GAMMA-CARBOXYGLUTAMIC ACID.  
 FT CARBOHYD 120 120 N-LINKED (GLCNAC. . .) (POTENTIAL).  
 FT CARBOHYD 144 144 N-LINKED (GLCNAC. . .) (POTENTIAL).  
 FT CARBOHYD 412 412 N-LINKED (GLCNAC. . .) (POTENTIAL).  
 FT CARBOHYD 552 552 N-LINKED (GLCNAC. . .) (POTENTIAL).  
 FT DISULFD 61 66 BY SIMILARITY.  
 FT DISULFD 91 104 BY SIMILARITY.  
 FT DISULFD 109 187 BY SIMILARITY.  
 FT DISULFD 130 170 BY SIMILARITY.  
 FT DISULFD 158 182 BY SIMILARITY.  
 FT DISULFD 215 292 BY SIMILARITY.  
 FT DISULFD 236 276 BY SIMILARITY.  
 FT DISULFD 264 287 BY SIMILARITY.  
 FT DISULFD 332 478 INTERCHAIN (BY SIMILARITY).  
 FT DISULFD 387 403 BY SIMILARITY.  
 FT DISULFD 532 546 BY SIMILARITY.  
 FT DISULFD 560 590 BY SIMILARITY.  
 SQ SEQUENCE 617 AA; 70411 MW; AD27DLB7144SDBD CRC64;

Query Match 43.9%; Score 86.5; DB 1; Length 617;  
 Best Local Similarity 42.2%; Pred. No. 6.2e-07;  
 Matches 19; Conservative 6; Mismatches 19; Indels 1; Gaps 1;

Oy 1 ANS-FLXXLRGSLXRXIXXICDPXKXIFEDVDLPLWRSKH 44  
 Db 44 ANSGFLBELRNGNLRECEVCSEAFEALESPDDIVEMAKY 88

RESULT 15  
 THRB\_MOUSE STANDARD; PRT: 618 AA.  
 AC P19221;  
 DT 01-NOV-1990 (Rel. 16, Created)  
 DT 01-NOV-1990 (Rel. 16, Last sequence update)  
 DT 15-JUN-2002 (Rel. 41, Last annotation update)  
 DE Prothrombin precursor (EC 3.4.21.5).  
 GN F2 OR CP2.  
 OS Mus musculus (Mouse).  
 OC Eukaryota; Metazoa; Chordata; Vertebrata; Euteleostomi;  
 OC Mammalia; Eutheria; Rodentia; Sciurognathi; Muridae; Mus.  
 NC NCBI\_TaxID=10090;  
 RN [1]  
 RP SEQUENCE FROM N.A.  
 RC STRAIN=C57BL/6; TISSUE=Liver;  
 RX MEDLINE=91023551; Pubmed=222810;  
 RA Fritzenr Degen S.J., Schaffner L.A., Jamison C.S., Grant S.G.,  
 RA Fitzgibbon J.J., Pai J.-A., Chapman V.M., Elliott R.W.;  
 RT "Characterization of the cDNA coding for mouse prothrombin and  
 RT localization of the gene on mouse chromosome 2.";  
 RL DNA Cell Biol. 9:487-498(1990).  
 RN [2]  
 RP SEQUENCE OF 384-618 FROM N.A.  
 RC TISSUE=Liver;  
 RX MEDLINE=92212913; Pubmed=1557383;  
 RA Banfield D.K., Macgillivray R.T.;  
 RT "Partial characterization of vertebrate prothrombin cDNAs:  
 RT amplification and sequence analysis of the B chain of thrombin from  
 RT nine different species.";  
 RL Proc. Natl. Acad. Sci. U.S.A. 89:2779-2783(1992).  
 CC -1- FUNCTION: THROMBIN, WHICH CLEAVES BONDS AFTER ANG & LYS, CONVERTS  
 CC FIBRINOGEN TO FIBRIN AND ACTIVATES FACTORS V, VII, VIII, XIII,  
 CC AND, IN COMPLEX WITH THROMBOMODULIN, PROTEIN C.  
 CC -1- CATALYTIC ACTIVITY: Preferential cleavage: Arg-I-Gly; activates  
 CC fibrinogen to fibrin and releases fibrinopeptide A and B.

CC -1- PTH: THE GAMMA-CARBOXYGLUTAMYL RESIDUES, WHICH BIND CALCIUM IONS,  
 CC RESULT FROM THE CARBOXYLATION OF GLUTAMYL RESIDUES BY A MICROSOMAL  
 CC ENZYME. THE VITAMIN K-DEPENDENT CARBOXYLASE. THE MODIFIED RESIDUES  
 CC ARE NECESSARY FOR THE CA-DEPENDENT INTERACTION WITH A NEGATIVELY  
 CC CHARGED PHOSPHOLIPID SURFACE, WHICH IS ESSENTIAL FOR THE CONVERSION  
 CC OF PROTHROMBIN TO THROMBIN.  
 CC -1- MISCELLANEOUS: PROTHROMBIN IS ACTIVATED ON THE SURFACE OF A  
 CC PHOSPHOLIPID MEMBRANE THAT BINDS THE AMINO END OF PROTHROMBIN &  
 CC FACTORS VA & XA IN CA-DEPENDENT INTERACTIONS; FACTOR XA REMOVES  
 CC THE ACTIVATION PEPTIDE & CLEAVES THE REMAINING PART INTO LIGHT &  
 CC HEAVY CHAINS. THE ACTIVATION PROCESS STARTS SLOWLY BECAUSE FACTOR  
 CC V ITSELF HAS TO BE ACTIVATED BY THE INITIAL, SMALL AMOUNTS OF  
 CC THROMBIN.  
 CC -1- MISCELLANEOUS: THROMBIN CAN ITSELF CLEAVE THE AMINO TERMINAL  
 CC FRAGMENT (FRAGMENT 1) OF THE PROTHROMBIN, PRIOR TO ITS ACTIVATION  
 CC BY FACTOR XA.  
 CC -1- SIMILARITY: BELONGS TO PEPTIDASE FAMILY S1.  
 CC -1- SIMILARITY: CONTRAINS 2 KRINGLE DOMAINS.  
 CC -----  
 CC This SWISS-PROT entry is copyright. It is produced through a collaboration  
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 CC or send an email to [license@isb-sib.ch](mailto:license@isb-sib.ch)).  
 CC -----  
 CC EMBL: X52308; CAA36548.1;  
 CC EMBL: M81394; AAA40435.1;  
 CC PIR: A35827; A35827.  
 CC HSSP: P00734; 1BPX.  
 CC MEROPS: S01.217; -.  
 CC MGD: MGI:88380; F2.  
 CC InterPro: IPR001314; Chymotrypsin.  
 CC InterPro: IPR002383; GLA blood.  
 CC InterPro: IPR000001; Kringle.  
 CC InterPro: IPR003966; Prothrombin.  
 CC InterPro: IPR001254; Ser-protease-try.  
 CC InterPro: IPR000294; VltK\_dep-GLA.  
 CC Pfam: PF00051; Kringle; 2.  
 CC Pfam: PF00089; trypsin; 1.  
 CC Pfam: PF00594; gla; 1.  
 CC PRINTS: PR00722; CHYMOTRYPSIN.  
 CC PRINTS: PR00001; GLABLOOD.  
 CC PRINTS: PR00018; KRINGLE.  
 CC PRINTS: PR01505; PROTHROMBIN.  
 CC ProDom: PD000395; Kringle; 2.  
 CC SMART: SM00069; GLA; 1.  
 CC SMART: SM00130; KR; 2.  
 CC SMART: SM00020; TRYP-spec; 1.  
 CC PROSITE: PS00011; GLU-CARBOXYLATION; 1.  
 CC PROSITE: PS00021; KRINGLE\_1; 2.  
 CC PROSITE: PS50070; KRINGLE\_2; 2.  
 CC PROSITE: PS50240; TRYPsin DOM; 1.  
 CC PROSITE: PS00134; TRYPsin\_HIS; 1.  
 CC PROSITE: PS00135; TRYPsin\_SER; 1.  
 CC Blood coagulation; Plasma; Calcium-binding; Glycoprotein; Repeat;  
 CC Vitamin K; zymogen; Gamma-carboxyglutamic acid; Acute phase; Liver;  
 CC Hydrolyase; Serine protease; Kringle; Signal.  
 CC SIGNAL 1 24  
 CC FT PROPEP 25 43  
 CC FT CHAIN 44 618  
 CC FT PEPTIDE 44 200  
 CC FT PEPTIDE 201 324  
 CC FT CHAIN 325 360  
 CC FT CHAIN 361 618  
 CC FT DOMAIN 109 187  
 CC FT DOMAIN 215 292  
 CC FT DOMAIN 361 618  
 CC FT SITE 200 201  
 CC FT SITE 324 325  
 CC FT SITE 360 361  
 CC ACT\_SITE 403 403  
 CC PROTHROMBIN.  
 CC ACTIVATION PEPTIDE (FRAGMENT 1).  
 CC ACTIVATION PEPTIDE (FRAGMENT 2).  
 CC THROMBIN LIGHT CHAIN (A).  
 CC THROMBIN HEAVY CHAIN (B).  
 CC KRINGLE 1.  
 CC KRINGLE 2.  
 CC SERINE PROTEASE.  
 CC CLEAVAGE (BY THROMBIN).  
 CC CLEAVAGE (BY FACTOR XA).  
 CC CLEAVAGE (BY FACTOR XA).  
 CC CHARGE RELAY SYSTEM (BY SIMILARITY).

```
FT ACT_SITE 459 459 CHARGE RELAY SYSTEM (BY SIMILARITY).
FT ACT_SITE 565 565 CHARGE RELAY SYSTEM (BY SIMILARITY).
FT MOD_RES 50 50 GAMMA-CARBOXYGLUTAMIC ACID.
FT MOD_RES 51 51 GAMMA-CARBOXYGLUTAMIC ACID.
FT MOD_RES 58 58 GAMMA-CARBOXYGLUTAMIC ACID.
FT MOD_RES 60 60 GAMMA-CARBOXYGLUTAMIC ACID.
FT MOD_RES 63 63 GAMMA-CARBOXYGLUTAMIC ACID.
FT MOD_RES 64 64 GAMMA-CARBOXYGLUTAMIC ACID.
FT MOD_RES 69 69 GAMMA-CARBOXYGLUTAMIC ACID.
FT MOD_RES 70 70 GAMMA-CARBOXYGLUTAMIC ACID.
FT MOD_RES 73 73 GAMMA-CARBOXYGLUTAMIC ACID.
FT MOD_RES 76 76 GAMMA-CARBOXYGLUTAMIC ACID.
FT MOD_RES 76 76 GAMMA-CARBOXYGLUTAMIC ACID.
FT DISULFID 61 66 BY SIMILARITY.
FT DISULFID 91 104 BY SIMILARITY.
FT DISULFID 109 187 BY SIMILARITY.
FT DISULFID 130 170 BY SIMILARITY.
FT DISULFID 158 182 BY SIMILARITY.
FT DISULFID 215 293 BY SIMILARITY.
FT DISULFID 236 276 BY SIMILARITY.
FT DISULFID 264 288 BY SIMILARITY.
FT DISULFID 333 479 INTERCHAIN (BY SIMILARITY).
FT DISULFID 388 404 BY SIMILARITY.
FT DISULFID 533 547 BY SIMILARITY.
FT DISULFID 561 591 BY SIMILARITY.
FT CARBOHYD 122 122 N-LINKED (GLCNAC. . .).
FT CARBOHYD 144 144 N-LINKED (GLCNAC. . .).
FT CARBOHYD 413 413 N-LINKED (GLCNAC. . .).
FT CARBOHYD 553 553 N-LINKED (GLCNAC. . .).
SQ SEQUENCE 618 AA: 70268 MM: B89F719AAFD601E0 CRC64;
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Query Match 43.9%; Score 86.5; DB 1; Length 618;  
Best Local Similarity 42.2%; Pred. No. 6.2e-07;  
Matches 19; Conservative 6; Mismatches 19; Indels 1; Gaps 1;

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OY 1 ANS-FLXXLRGSLRXKXICXIXICDFXXAKXIFEDVDGTLAFWSKH 44
DB 44 ANSGFLLELRKGNLERECVEQSYEEAFEALESPODTPVFWAKY 88
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Search completed: May 16, 2003, 10:14:50  
Job time : 12 secs

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GenCore version 5.1.4-P5.4578  
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OM protein - protein search, using sw model

Run on: May 16, 2003, 10:12:19 ; Search time 29 Seconds  
(without alignments)  
312.623 Million cell updates/sec

Title: SEQ1-4EDITS  
Perfect score: 197  
Sequence: 1 ANSFLXLRQGSIXRXCIXX.....XXAKXIFEDVDDTLAFWSKH 44

Scoring table: BLOSUM62  
Gapop 10.0 , Gapext 0.5

Searched: 671580 seqs, 206047115 residues

Total: number of hits satisfying chosen parameters: 671580

Minimum DB seq length: 0  
Maximum DB seq length: 2000000000

Post-processing: Minimum Match 0%  
Maximum Match 100%  
Listing first 45 summaries

Database :

1: SP\_ARCHAEA:\*  
2: SP\_BACTERIA:\*  
3: SP\_FUNGI:\*  
4: SP\_HUMAN:\*  
5: SP\_INVERTEBRATE:\*  
6: SP\_MAMMAL:\*  
7: SP\_MHC:\*  
8: SP\_ORGANELLE:\*  
9: SP\_PHAGE:\*  
10: SP\_PLANT:\*  
11: SP RODENT:\*  
12: SP\_VIRUS:\*  
13: SP\_VERTEBRATE:\*  
14: SP\_UNCLASSIFIED:\*  
15: SP\_VIRUS:\*  
16: SP\_BACTERIAP:\*  
17: SP\_ARCHAEP:\*

Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

# SUMMARIES

Result No.	Score	Query Match	Length	DB ID	Description
1	151	76.6	456	6 Q9TRRO	Q9TRRO canis famli
2	140	71.1	460	11 Q91WN8	Q91WN8 mus musculu
3	134	68.0	460	11 Q99PC6	Q99PC6 mus musculu
4	115	58.4	482	11 Q63207	Q63207 rattus norv
5	101	51.3	481	11 Q54740	Q54740 mus musculu
6	101	51.3	481	11 Q99L32	Q99L32 mus musculu
7	101	51.3	481	11 Q88947	Q88947 mus musculu
8	99	50.3	701	4 Q96F08	Q96F08 homo sapien
9	95	48.2	469	6 Q9GMD9	Q9GMD9 ornithorhyn
10	85	43.1	650	4 Q9NSD0	Q9NSD0 homo sapien
11	85	43.1	650	4 Q16519	Q16519 homo sapien
12	84	42.6	100	4 Q15253	Q15253 homo sapien
13	82.5	41.9	542	5 Q8TF13	Q8TF13 halocynthia
14	80	40.6	446	11 Q61109	Q61109 mus musculu
15	80	40.6	456	6 Q14316	Q14316 homo sapien
16	80	40.6	461	6 Q95ND7	Q95ND7 pan troglod

17	80	40.6	461	6 Q95ND6	Q95ND6 pan troglod
18	78	39.6	138	6 Q28994	Q28994 sus scrofa
19	78	39.6	607	13 Q91001	Q91001 gallus gall
20	78	39.6	648	6 Q29094	Q29094 sus scrofa
21	73.5	37.3	433	13 Q90YK1	Q90YK1 brachydanio
22	73	37.1	49	6 Q95ME8	Q95ME8 bos taurus
23	73	37.1	399	11 Q9COW3	Q9COW3 mus musculu
24	72	36.5	98	13 P82807	P82807 neochlis sc
25	72	36.5	608	13 Q9PTW7	Q9PTW7 struthio ca
26	66	33.5	25	11 Q9QVH6	Q9QVH6 rattus sp.
27	65	33.0	179	4 Q8TAS3	Q8TAS3 homo sapien
28	65	33.0	198	11 Q8R182	Q8R182 mus musculu
29	65	33.0	673	11 Q61592	Q61592 mus musculu
30	65	33.0	674	11 Q99K57	Q99K57 mus musculu
31	64	32.5	674	11 Q63772	Q63772 rattus sp.
32	63	32.0	678	4 Q14393	Q14393 homo sapien
33	56.5	28.7	459	10 Q9SE22	Q9SE22 oryza sativ
34	56.5	28.7	606	10 Q95JG9	Q95JG9 arabidopsis
35	56.5	28.7	651	10 Q85218	Q85218 oryza sativ
36	55.5	28.2	575	10 Q94E17	Q94E17 oryza sativ
37	54.5	27.7	567	10 Q8W4J2	Q8W4J2 arabidopsis
38	54.5	27.7	603	10 Q9LPG7	Q9LPG7 arabidopsis
39	53.5	27.2	196	10 Q04284	Q04284 selaginella
40	53.5	27.2	431	10 Q94EY5	Q94EY5 arabidopsis
41	53.5	27.2	506	10 Q9SPF0	Q9SPF0 oryza sativ
42	53.5	27.2	506	10 Q9SE23	Q9SE23 oryza sativ
43	53.5	27.2	543	10 Q9MB23	Q9MB23 arabidopsis
44	53.5	27.2	568	10 Q9ASC3	Q9ASC3 oryza sativ
45	53.5	27.2	576	10 Q9C9U4	Q9C9U4 arabidopsis

## ALIGNMENTS

RESULT 1	ID	Q9TRRO	PRELIMINARY;	PRT;	456 AA.
AC	Q9TRRO				
DT	01-MAY-2000 (TREMBLrel. 13, Created)				
DT	01-MAY-2000 (TREMBLrel. 13, Last sequence update)				
DE	01-MAR-2002 (TREMBLrel. 20, Last annotation update)				
DE	Protein C precursor.				
GN	PROC.				
OS	Canis familiaris (Dog).				
OC	Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;				
OC	Mammalia; Eutheria; Carnivora; Fissipedia; Canidae; Canis.				
OX	NCBI_TaxID=9615;				
RN	[1]				
RP	SEQUENCE FROM N.A.				
RA	Leeb T., Kopp T., Deppe A., Breen M., Matlis U., Brunnberg L.,				
RA	Brenig B.;				
RT	"Molecular characterization and chromosomal assignment of the canine				
RT	protein C gene."				
RL	Mamm. Genome 10:135-139(1999).				
RN	[2]				
RP	SEQUENCE FROM N.A.				
RX	MEDLINE-99371952; PubMed-10443005;				
RA	Leeb T., Pfeiffer I., Kopp T., Deppe A., Brenig B.;				
RT	"Analysis of canine protein C gene polymorphisms."				
RL	Anim. Genet. 30:237-238(1999).				
CC	-1- SIMILARITY: BELONGS TO PEPTIDASE FAMILY S1; ALSO KNOWN AS THE				
CC	TRYPSIN FAMILY.				
DR	EMBL: AJ001979; CAA05126.1; -				
DR	HSSP: P04070; 1PCU.				
DR	MEROPS: S01.218; -				
DR	Interpro: IPR000152; Asx_hydroxyl.				
DR	Interpro: IPR001314; Chymotrypsin.				
DR	Interpro: IPR000561; EGF-like.				
DR	Interpro: IPR001881; EGF Ca.				
DR	Interpro: IPR002383; GLA_blood.				
DR	Interpro: IPR001254; Ser_protease_Try.				
DR	Interpro: IPR000294; Vitk_dep_GLA.				
DR	Pfam: PF00008; EGF; 2.				

DR Pfam: PF00594; gla; 1.  
 DR Pfam: PF00089; trypsin; 1.  
 DR PRINTS; PR00722; CHYMOTRYPSIN.  
 DR PRINTS; PR00001; GLABLOOD.  
 DR SMART; SM00181; EGF; 2.  
 DR SMART; SM00069; gla; 1.  
 DR SMART; SM00020; TRYP\_SPE; 1.  
 DR PROSITE; PS00010; ASX\_HYDROXYL; 1.  
 DR PROSITE; PS00022; EGF\_1; UNKNOWN\_1.  
 DR PROSITE; PS01186; EGF\_2; 2.  
 DR PROSITE; PS01187; EGF\_CA; 1.  
 DR PROSITE; PS00011; GLU\_CARBOXYLATION; 1.  
 DR PROSITE; PS00240; TRYPsin.DOM; 1.  
 DR PROSITE; PS00134; TRYPsin.HIS; UNKNOWN\_1.  
 DR PROSITE; PS00135; TRYPsin.SER; 1.  
 DR Calcium-binding; EGF-like domain; Glycoprotein; Hydrolase;  
 KM Hydroxylation; Repeat; Serine protease; Signal.  
 FT SIGNAL 1 42  
 FT CHAIN 43 192 POTENTIAL.  
 FT CHAIN 193 194 PROTEIN C CONNECTING DIPEPTIDE.  
 FT CHAIN 195 456 PROTEIN C HEAVY CHAIN.  
 SQ SEQUENCE 456 AA; 50813 MW; 7AD3AB0C134E59FF CRC64;

Query Match 76.6%; Score 151; DB 6; Length 456;  
 Best Local Similarity 63.6%; Pred. No. 1.6e-17;  
 Matches 28; Conservative 6; Mismatches 10; Indels 0; Gaps 0;

OY 1 ANSFLXLLRQSLXRXICIXICDFFXXAKXIFEDVDTLAFWSKH 44  
 DB 43 ANSFLLEIRAGSLERECMEICDFFERAKEIFQNVDDTLAFWSKY 86

RESULT 2  
 ID 091WN8 PRELIMINARY; PRT; 460 AA;  
 AC 091WN8;  
 DT 01-DEC-2001 (TREMBLrel. 19, Created)  
 DT 01-DEC-2001 (TREMBLrel. 19, last sequence update)  
 DT 01-JUN-2002 (TREMBLrel. 21, last annotation update)  
 DE Similar to protein C.  
 GN PROC.  
 OS Mus musculus (Mouse).  
 OC Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
 OC Mammalia; Eutheria; Rodentia; Sciurognathi; Muridae; Murinae; Mus.  
 CX NCBI\_TaxID=10090;  
 RN [1]  
 RP SEQUENCE FROM N.A.  
 RC TISSUE=LIVER;  
 RA Strausberg R.;  
 RL Submitted (SEP-2001) to the EMBL/Genbank/DBJ databases.  
 DR MGD; MGI:97771; PROC.  
 DR InterPro; IPR000152; Asx\_hydroxyl.  
 DR InterPro; IPR000561; EGF-like.  
 DR InterPro; IPR001881; EGF\_CA.  
 DR InterPro; IPR001254; Ser\_protease\_Try.  
 DR InterPro; IPR000294; VitK\_dep\_GLA.  
 DR Pfam; PF00008; EGF; 2.  
 DR Pfam; PF00594; gla; 1.  
 DR Pfam; PF00089; trypsin; 1.  
 DR PROSITE; PS00010; ASX\_HYDROXYL; UNKNOWN\_1.  
 DR PROSITE; PS00022; EGF\_1; UNKNOWN\_1.  
 DR PROSITE; PS01186; EGF\_2; UNKNOWN\_2.  
 DR PROSITE; PS01187; EGF\_CA; UNKNOWN\_1.  
 DR PROSITE; PS00011; GLU\_CARBOXYLATION; UNKNOWN\_1.  
 DR PROSITE; PS00240; TRYPsin.DOM; 1.  
 DR PROSITE; PS00134; TRYPsin.HIS; UNKNOWN\_1.  
 DR PROSITE; PS00135; TRYPsin.SER; UNKNOWN\_1.  
 KM Hydrolase; Serine protease.  
 SQ SEQUENCE 460 AA; 51818 MW; 0117F26E68FCC274 CRC64;

Query Match 71.1%; Score 140; DB 11; Length 460;  
 Best Local Similarity 59.1%; Pred. No. 1.3e-15;

Matches 26; Conservative 7; Mismatches 11; Indels 0; Gaps 0;  
 OY 1 ANSFLXLLRQSLXRXICIXICDFFXXAKXIFEDVDTLAFWSKH 44  
 DB 42 ANSFLLEIRAGSLERECMEICDFFERAKEIFQNVDDTLAFWIKY 85

RESULT 3  
 ID 099PC6 PRELIMINARY; PRT; 460 AA.  
 AC 099PC6;  
 DT 01-JUN-2001 (TREMBLrel. 17, Created)  
 DT 01-JUN-2001 (TREMBLrel. 17, last sequence update)  
 DT 01-JUN-2002 (TREMBLrel. 21, last annotation update)  
 DE Anticoagulant protein C.  
 GN PROC.  
 OS Mus musculus (Mouse).  
 OC Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
 OC Mammalia; Eutheria; Rodentia; Sciurognathi; Muridae; Murinae; Mus.  
 CX NCBI\_TaxID=10090;  
 RN [1]  
 RP SEQUENCE FROM N.A.  
 RC STRAIN=C57BL;  
 RA Korf I.;  
 RL "Complete sequence of UC72A01."  
 RL Submitted (NOV-2000) to the EMBL/Genbank/DBJ databases.  
 CC -I- SIMILARITY: BELONGS TO PEPTIDASE FAMILY S1; ALSO KNOWN AS THE  
 CC TRYPSIN FAMILY.  
 CC EMBL; AF318182; AAK07918.1; -  
 DR HSSP; P04070; 1PCU.  
 DR MEROPS; S01\_218; -  
 DR MGD; MGI:97771; PROC.  
 DR InterPro; IPR000152; Asx\_hydroxyl.  
 DR InterPro; IPR001314; Chymotrypsin.  
 DR InterPro; IPR000561; EGF-like.  
 DR InterPro; IPR001881; EGF\_CA.  
 DR InterPro; IPR002383; GLA\_blood.  
 DR InterPro; IPR001254; Ser\_protease\_Try.  
 DR InterPro; IPR000294; VitK\_dep\_GLA.  
 DR Pfam; PF00008; EGF; 2.  
 DR Pfam; PF00594; gla; 1.  
 DR Pfam; PF00089; trypsin; 1.  
 DR PRINTS; PR00722; CHYMOTRYPSIN.  
 DR PRINTS; PR00001; GLABLOOD.  
 DR SMART; SM00001; EGF; 2.  
 DR SMART; SM00001; EGF\_2.  
 DR SMART; SM00069; gla; 1.  
 DR SMART; SM00020; TRYP\_SPE; 1.  
 DR PROSITE; PS00010; ASX\_HYDROXYL; 1.  
 DR PROSITE; PS00022; EGF\_1; UNKNOWN\_1.  
 DR PROSITE; PS01186; EGF\_2; 2.  
 DR PROSITE; PS01187; EGF\_CA; 1.  
 DR PROSITE; PS00011; GLU\_CARBOXYLATION; 1.  
 DR PROSITE; PS00240; TRYPsin.DOM; 1.  
 DR PROSITE; PS00134; TRYPsin.HIS; UNKNOWN\_1.  
 DR PROSITE; PS00135; TRYPsin.SER; 1.  
 KM Calcium-binding; EGF-like domain; Glycoprotein; Hydrolase;  
 KM Hydroxylation; Repeat; Serine protease.  
 SQ SEQUENCE 460 AA; 51784 MW; 0293BC25E9D3ED16 CRC64;

Query Match 68.0%; Score 134; DB 11; Length 460;  
 Best Local Similarity 56.8%; Pred. No. 1.4e-14;  
 Matches 25; Conservative 7; Mismatches 12; Indels 0; Gaps 0;

OY 1 ANSFLXLLRQSLXRXICIXICDFFXXAKXIFEDVDTLAFWSKH 44  
 DB 42 ANSFLLEIRAGSLERECMEICDFFERAKEIFQNVDDTLAFWIKY 85

RESULT 4  
 ID 063207 PRELIMINARY; PRT; 482 AA.  
 AC 063207;

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DT 01-NOV-1996 (Tremblrel. 01, Created)
DT 01-NOV-1996 (Tremblrel. 01, Last sequence update)
DT 01-MAR-2002 (Tremblrel. 20, Last annotation update)
DE Factor X.
OS Rattus norvegicus (Rat).
OC Eukaryota; Metazoa; Chordata; Craniala; Vertebrata; Euteleostomi;
OC Mammalia; Eutheria; Rodentia; Sciurognathi; Muridae; Murinae; Rattus.
NCBI_TaxID=10116;
RN [1]
RP SEQUENCE FROM N.A.
RC STRAIN=SPRAGUE-DAWLEY;
RX MEDLINE=96093366; PubMed=8578539;
Stanton C., Ross R.P., Hutson S., Wallin R.;
RT "Evidence for competition between vitamin K-dependent clotting factors
for intracellular processing by the vitamin K-dependent gamma-
carboxylase."
RT Thromb. Res. 80:63-73(1995).
CC -1- SIMILARITY: BELONGS TO PEPTIDASE FAMILY S1; ALSO KNOWN AS THE
TRYPSIN FAMILY.
CC EMBL: X79807; CAA56202.1; -.
DR HSSP: P00742; 1XKA.
DR MEROPS: S01.216; -.
DR InterPro: IPR000152; Asx_hydroxyl.
DR InterPro: IPR001314; Chymotrypsin.
DR InterPro: IPR000561; EGF_1like.
DR InterPro: IPR000742; EGF_2.
DR InterPro: IPR001881; EGF_Ca.
DR InterPro: IPR002383; GLA_blood.
DR InterPro: IPR001254; Ser_protease_Try.
DR InterPro: IPR000294; Vitk_dep_GLA.
DR Pfam: PF000594; g1a; 1.
DR Pfam: PF00089; trypsin; 1.
DR PRINTS: PR00089; CHYMOTRYPSIN.
DR PRINTS: PR00001; GLABLOOD.
DR SMART: SM00179; EGF_CA; 1.
DR SMART: SM00001; EGF_1like; 1.
DR SMART: SM00069; GLA; 1.
DR SMART: SM00020; TRYP_SPE; 1.
DR PROSITE: PS00010; ASX_HYDROXYL; 1.
DR PROSITE: PS00022; EGF_1; UNKNOWN_1.
DR PROSITE: PS01186; EGF_2; 2.
DR PROSITE: PS01187; EGF_CA; 1.
DR PROSITE: PS00011; GLU_CARBOXYLATION; 1.
DR PROSITE: PS02040; TRYPSIN_DOM; 1.
DR PROSITE: PS00134; TRYPSIN_HTS; UNKNOWN_1.
DR PROSITE: PS00135; TRYPSIN_SER; 1.
KW Calcium-binding; EGF-like domain; Glycoprotein; Hydrolase;
KW Hydroxylation; Repeat; Serine protease.
SQ SEQUENCE 482 AA; 54265 MW; 0284678E3954A698 CRC64;

Query Match 58.4%; Score 115; DB 11; Length 482;
Best Local Similarity 43.2%; Pred. No. 2.8e-11;
Matches 19; Conservative 10; Mismatches 15; Indels 0; Gaps 0;

QY 1 ANSFLXLRGSLXKRCIXXICDFFXAKXIFEDVDTLAFWSKH 44
DB 41 ANSFEERKGNLERECMEICSEFEARVFEDEKTEKRWTKY 84

RESULT 5
ID 054740 PRELIMINARY; PRT; 481 AA.
AC 054740;
DT 01-JUN-1998 (Tremblrel. 06, Created)
DT 01-JUN-1998 (Tremblrel. 06, Last sequence update)
DT 01-JUN-2002 (Tremblrel. 21, Last annotation update)
DE Coagulation factor X precursor (EC 3.4.21.6).
GN F10 OR F10.
OS Mus musculus (Mouse).
OC Plasmid pBluescript.
OC Eukaryota; Metazoa; Chordata; Craniala; Vertebrata; Euteleostomi;
OC Mammalia; Eutheria; Rodentia; Sciurognathi; Muridae; Murinae; Mus.

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OX NCBI_TaxID=10090;
RN [1]
RP SEQUENCE FROM N.A.
RC TISSUE=LIVER;
RX MEDLINE=98454993; PubMed=9783672;
Heidmann H.H., Kontemann R.E.;
RT "Cloning and recombinant expression of mouse coagulation factor X."
RT Thromb. Res. 92:33-41(1998).
CC -1- SIMILARITY: BELONGS TO PEPTIDASE FAMILY S1; ALSO KNOWN AS THE
TRYPSIN FAMILY.
CC EMBL: A022677; CAA10933.1; -.
DR HSSP: P00742; 1XKA.
DR MEROPS: S01.216; -.
DR MGD: MGI:103107; F10.
DR InterPro: IPR000152; Asx_hydroxyl.
DR InterPro: IPR001314; Chymotrypsin.
DR InterPro: IPR000561; EGF_1like.
DR InterPro: IPR000742; EGF_2.
DR InterPro: IPR001881; EGF_Ca.
DR InterPro: IPR002383; GLA_blood.
DR InterPro: IPR001254; Ser_protease_Try.
DR InterPro: IPR000294; Vitk_dep_GLA.
DR Pfam: PF00008; EGF_2.
DR Pfam: PF00594; g1a; 1.
DR Pfam: PF00089; trypsin; 1.
DR PRINTS: PR00722; CHYMOTRYPSIN.
DR PRINTS: PR00001; GLABLOOD.
DR SMART: SM00179; EGF_CA; 1.
DR SMART: SM00001; EGF_1like; 1.
DR SMART: SM00069; GLA; 1.
DR SMART: SM00020; TRYP_SPE; 1.
DR PROSITE: PS00010; ASX_HYDROXYL; UNKNOWN_1.
DR PROSITE: PS00022; EGF_1; UNKNOWN_1.
DR PROSITE: PS01186; EGF_2; 2.
DR PROSITE: PS01187; EGF_CA; 1.
DR PROSITE: PS00011; GLU_CARBOXYLATION; 1.
DR PROSITE: PS02040; TRYPSIN_DOM; 1.
DR PROSITE: PS00134; TRYPSIN_HTS; UNKNOWN_1.
DR PROSITE: PS00135; TRYPSIN_SER; 1.
KW Calcium-binding; EGF-like domain; Glycoprotein; Hydrolase; Plasmid;
KW Repeat; Serine protease; Signal.
FT SIGNAL 1 40
FT CHAIN 41 481 COAGULATION FACTOR X.
SQ SEQUENCE 481 AA; 53986 MW; CF702DE5EF9D97AE CRC64;

Query Match 51.3%; Score 101; DB 11; Length 481;
Best Local Similarity 38.0%; Pred. No. 7.4e-09;
Matches 17; Conservative 10; Mismatches 17; Indels 0; Gaps 0;

QY 1 ANSFLXLRGSLXKRCIXXICDFFXAKXIFEDVDTLAFWSKH 44
DB 41 ANSFEERKGNLERECMEICSEFEARVFEDEKTEKRWTKY 84

RESULT 6
ID 099132 PRELIMINARY; PRT; 481 AA.
AC 099132;
DT 01-JUN-2001 (Tremblrel. 17, Created)
DT 01-JUN-2001 (Tremblrel. 17, Last sequence update)
DT 01-JUN-2002 (Tremblrel. 21, Last annotation update)
DE Coagulation factor X.
GN F10.
OS Mus musculus (Mouse).
OC Eukaryota; Metazoa; Chordata; Craniala; Vertebrata; Euteleostomi;
OC Mammalia; Eutheria; Rodentia; Sciurognathi; Muridae; Murinae; Mus.
NCBI_TaxID=10090;
RN [1]
RP SEQUENCE FROM N.A.
RC Strausberg R.;
RX Submitted (Feb-2001) to the EMBL/GenBank/DBJ databases.
RT -1- SIMILARITY: BELONGS TO PEPTIDASE FAMILY S1; ALSO KNOWN AS THE
TRYPSIN FAMILY.

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DR EMBL: BC003877; AA03877.1; -.
DR HSSP: P00742; 1XKA.
DR MEROPS: S01.216; -.
DR MGD: MGI:103107; F10.
DR InterPro: IPR000152; Asx_hydroxyl.
DR InterPro: IPR001314; Chymotrypsin.
DR InterPro: IPR000561; EGF-like.
DR InterPro: IPR000742; EGF_2.
DR InterPro: IPR001881; EGF_Ca.
DR InterPro: IPR001438; EGF_11.
DR InterPro: IPR002383; GLA_blood.
DR InterPro: IPR001254; Ser_protease_Try.
DR InterPro: IPR000294; VitK_dep_GLA.
DR Pfam: PF00008; EGF_2.
DR Pfam: PF00594; gla; 1.
DR Pfam: PF00089; trypsin; 1.
DR PRINTS: PR00722; CHYMOTRYPSIN.
DR PRINTS: PR00010; EGF_BLOOD.
DR PRINTS: PR00001; GLABLOOD.
DR SMART: SM00001; EGF_1like; 2.
DR SMART: SM00001; EGF_1like; 2.
DR SMART: SM00069; GLA; 1.
DR SMART: SM00020; TRYP_SPE; 1.
DR PROSITE: PS00010; ASX_HYDROXYL; UNKNOWN_1.
DR PROSITE: PS00022; EGF_1; UNKNOWN_1.
DR PROSITE: PS01186; EGF_2; 2.
DR PROSITE: PS01187; EGF_CA; 1.
DR PROSITE: PS00011; GLU_CARBOXYLATION; 1.
DR PROSITE: PS00240; TRYPsin_DOM; 1.
DR PROSITE: PS00134; TRYPsin_HIS; UNKNOWN_1.
DR PROSITE: PS00135; TRYPsin_SER; 1.
KW Calcium-binding; EGF-like domain; Glycoprotein; Hydrolase; Repeat;
KW Serine protease.
SQ SEQUENCE 481 AA; 54004 MW; BD88E96C8A0B7E7F CRC64;

Query Match          51.3%; Score 101; DB 11; Length 481;
Best Local Similarity 38.6%; Pred. No. 7.4e-09;
Matches 17; Conservative 10; Mismatches 17; Indels 0; Gaps 0;

Oy 1 ANSFLXLRQGSIXKXCIXICDFYXAKXIFEDVDTLAFNSKH 44
    |||| :||:| | | | | : |||| : | : ||:
Db 41 ANSFEEFKGNLRECKMEICSYEVRIFEDDEKTEKWTYKY 84

RESULT 7
ID 088947 PRELIMINARY; PRT; 481 AA.
AC 088947;
DT 01-NOV-1998 (TREMblrel. 08, Created)
DT 01-NOV-1998 (TREMblrel. 08, Last sequence update)
DT 01-MAR-2002 (TREMblrel. 20, Last annotation update)
DE Coagulation factor X precursor.
GN F10.
OS Mus musculus (Mouse).
OC Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC Mammalia; Eutheria; Rodentia; Sciurognathi; Muridae; Murinae; Mus.
OX NCBI_TaxID=10090;
RN [1]
RP SEQUENCE FROM N.A.
RC STRAIN=C57BL6 X CBA; TISSUE=LIVER;
RX MEDLINE=98347933; PubMed=9684791;
RA Liang Z., Cooper A., DeFord M.E., Carmeliet P., Collen D.,
RA Castellino F.J., Rosen E.D.,
RT "Cloning and characterization of a cDNA encoding murine coagulation
RT factor X.";
RL Thromb. Haemost. 80:87-91(1998).
RN [2]
RP SEQUENCE FROM N.A.
RC STRAIN=129SJJ;
RA Cooper A., Liang Z., Castellino F.J., Rosen E.D.;
RT "Cloning and Characterization of the Murine Factor X Gene.";
RL Thromb. Haemost. 0:0-0(2000).
CC -1- SIMILARITY: BELONGS TO PEPTIDASE FAMILY S1; ALSO KNOWN AS THE

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CC TRYPSIN FAMILY.
DR EMBL: AF087644; AAC36345.1; -.
DR EMBL: AF211347; AAF22980.1; -.
DR HSSP: P00742; 1XKA.
DR MEROPS: S01.216; -.
DR MGD: MGI:103107; F10.
DR InterPro: IPR000152; Asx_hydroxyl.
DR InterPro: IPR001314; Chymotrypsin.
DR InterPro: IPR000561; EGF-like.
DR InterPro: IPR000742; EGF_2.
DR InterPro: IPR001881; EGF_Ca.
DR InterPro: IPR002383; GLA_blood.
DR InterPro: IPR001254; Ser_protease_Try.
DR InterPro: IPR000294; VitK_dep_GLA.
DR Pfam: PF00008; EGF_2.
DR Pfam: PF00594; gla; 1.
DR Pfam: PF00089; trypsin; 1.
DR PRINTS: PR00722; CHYMOTRYPSIN.
DR PRINTS: PR00001; GLABLOOD.
DR SMART: SM00179; EGF_CA; 1.
DR SMART: SM00001; EGF_1like; 1.
DR SMART: SM00069; GLA; 1.
DR SMART: SM00020; TRYP_SPE; 1.
DR PROSITE: PS00010; ASX_HYDROXYL; UNKNOWN_1.
DR PROSITE: PS00022; EGF_1; UNKNOWN_1.
DR PROSITE: PS01186; EGF_2; 2.
DR PROSITE: PS01187; EGF_CA; 1.
DR PROSITE: PS00011; GLU_CARBOXYLATION; 1.
DR PROSITE: PS00240; TRYPsin_DOM; 1.
DR PROSITE: PS00134; TRYPsin_HIS; UNKNOWN_1.
DR PROSITE: PS00135; TRYPsin_SER; 1.
KW Calcium-binding; EGF-like domain; Glycoprotein; Hydrolase; Repeat;
KW Serine protease; Signal.
FT SIGNAL 1 40
FT CHAIN 41 481
SQ SEQUENCE 481 AA; 54018 MW; 8AC09DE5E9D271E CRC64;

Query Match          51.3%; Score 101; DB 11; Length 481;
Best Local Similarity 38.6%; Pred. No. 7.4e-09;
Matches 17; Conservative 10; Mismatches 17; Indels 0; Gaps 0;

Oy 1 ANSFLXLRQGSIXKXCIXICDFYXAKXIFEDVDTLAFNSKH 44
    |||| :||:| | | | | : |||| : | : ||:
Db 41 ANSFEEFKGNLRECKMEICSYEVRIFEDDEKTEKWTYKY 84

RESULT 8
ID 096P08 PRELIMINARY; PRT; 701 AA.
AC 096P08;
DT 01-DEC-2001 (TREMblrel. 19, Created)
DT 01-DEC-2001 (TREMblrel. 19, Last sequence update)
DT 01-MAR-2002 (TREMblrel. 20, Last annotation update)
DE Factor VII active site mutant immunconjugate.
GN Homo sapiens (Human).
OS Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC Mammalia; Eutheria; Primates; Catarrhini; Homnidae; Homo.
OX NCBI_TaxID=9606;
RN [1]
RP SEQUENCE FROM N.A.
RC MEDLINE=21477448; PubMed=11593034;
RX Hu Z., Garen A.;
RT "Targeting tissue factor on tumor vascular endothelial cells and tumor
RT cells for immunotherapy in mouse models of prostatic cancer.";
RL Proc. Natl. Acad. Sci. U.S.A. 98:12180-12185(2001).
DR EMBL: AF272774; AAK58686.1; -.
DR InterPro: IPR000152; Asx_hydroxyl.
DR InterPro: IPR000561; EGF-like.
DR InterPro: IPR000742; EGF_2.
DR InterPro: IPR001881; EGF_CA.
DR InterPro: IPR003006; Ig_MHC.
DR InterPro: IPR001254; Ser_protease_Try.
DR InterPro: IPR000294; VitK_dep_GLA.

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DR Pfam: PF00008; EGF_2.
DR Pfam: PF00594; g1a; 1.
DR Pfam: PF00047; 1g; 2.
DR Pfam: PF00089; trypsin; 1.
DR SMART: SM00181; EGF_2.
DR PROSITE: PS00010; ASX_HYDROXYL; UNKNOWN_1.
DR PROSITE: PS00022; EGF_1; UNKNOWN_1.
DR PROSITE: PS01186; EGF_2; UNKNOWN_1.
DR PROSITE: PS01187; EGF_CA; UNKNOWN_1.
DR PROSITE: PS00011; GLU_CARBOXYLATION; UNKNOWN_1.
DR PROSITE: PS00290; IG_HMC; UNKNOWN_1.
DR PROSITE: PS00240; TRYPSIN_DOM; 1.
DR PROSITE: PS00134; TRYPSIN_HIS; UNKNOWN_1.
DR PROSITE: PS00135; TRYPSIN_SER; UNKNOWN_1.
DR Hydrolase: Serine protease.
SQ SEQUENCE 701 AA; 77826 MW; 94AC6CEB42CC992F CRC64;

Query Match          50.3%; Score 99; DB 4; Length 701;
Best Local Similarity 48.8%; Pred. No. 2.5e-08;
Matches 20; Conservative 4; Mismatches 17; Indels 0; Gaps 0;

OY 1 ANSFLXLRGSLXRXIXICDFXAKXIFEDVDTLAW 41
Db 61 ANAFLELRPSLEKECEKCSFEARELFFKDAERTKLFW 101

RESULT 9
O9GMD9 PRELIMINARY; PRT; 469 AA.
AC O9GMD9;
DT 01-MAR-2001 (Tremblrel. 16, Created)
DT 01-MAR-2001 (Tremblrel. 16, Last sequence update)
DT 01-JUN-2002 (Tremblrel. 21, Last annotation update)
DE Coagulation factor X.
OS Ornithorhynchus anatinus (Duckbill platypus).
OC Eukaryota; Metazoa; Chordata; Craniala; Vertebrata; Euteleostomi;
OC Mammalia; Monotremata; Ornithorhynchidae; Ornithorhynchus.
OX NCBI_TaxID=9258;
RN (1)
RP SEQUENCE FROM N.A.
RX MEDLINE=21015017; Pubmed=1132153;
RA Poorefishar M., Aveskogh M., Munday B., Hellman L.;
RT Identification and structural analysis of four serine proteases in a
monotreme, the platypus, Ornithorhynchus anatinus."
RL Immunogenetics 52:19-28(2000).
CC -1- SIMILARITY: BELONGS TO PEPTIDASE FAMILY S1; ALSO KNOWN AS THE
CC TRYPSIN FAMILY.
DR EMBL: AF275654; ANG00453.1; -
DR HSP: P00742; 1XKB.
DR MEROPS: S01.216; -
DR InterPro: IPR000152; ASX_hydroxyl.
DR InterPro: IPR001314; Chymotrypsin.
DR InterPro: IPR000561; EGF-like.
DR InterPro: IPR000742; EGF_2.
DR InterPro: IPR001881; EGF_CA.
DR InterPro: IPR002383; GLA_blood.
DR InterPro: IPR001254; Ser.protease_Try.
DR InterPro: IPR000294; VitK_dep_GLA.
DR Pfam: PF00008; EGF_2.
DR Pfam: PF00594; g1a; 1.
DR Pfam: PF00089; trypsin; 1.
DR PRINTS: PR00722; CHYMOTRYPSIN.
DR PRINTS: PR00001; GLABLOOD.
DR SMART: SM00181; EGF_2.
DR SMART: SM00179; EGF_CA; 1.
DR SMART: SM00001; EGF_Like; 2.
DR SMART: SM00069; GLA; 1.
DR SMART: SM00020; TRYP_SPC; 1.
DR PROSITE: PS00010; ASX_HYDROXYL; UNKNOWN_1.
DR PROSITE: PS00022; EGF_1; UNKNOWN_1.
DR PROSITE: PS01186; EGF_2; 2.
DR PROSITE: PS01187; EGF_CA; 1.
DR PROSITE: PS00011; GLU_CARBOXYLATION; 1.

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DR PROSITE: PS00240; TRYPSIN_DOM; 1.
DR PROSITE: PS00134; TRYPSIN_HIS; UNKNOWN_1.
DR PROSITE: PS00135; TRYPSIN_SER; 1.
DR Hydrolase: Serine protease.
SQ SEQUENCE 469 AA; 52196 MW; 4C66C230D0758F6A CRC64;

Query Match          48.2%; Score 95; DB 6; Length 469;
Best Local Similarity 40.5%; Pred. No. 7.9e-08;
Matches 17; Conservative 7; Mismatches 18; Indels 0; Gaps 0;

OY 1 ANSFLXLRGSLXRXIXICDFXAKXIFEDVDTLAWFS 42
Db 41 ANSLFEELKGNLERECNEETCSYEAREVVEDDTKNEFWN 82

RESULT 10
O9NSD0 PRELIMINARY; PRT; 650 AA.
AC O9NSD0;
DT 01-OCT-2000 (Tremblrel. 15, Created)
DT 01-OCT-2000 (Tremblrel. 15, Last sequence update)
DT 01-JUN-2002 (Tremblrel. 21, Last annotation update)
DE Protein S precursor.
OS Homo sapiens (Human).
OC Eukaryota; Metazoa; Chordata; Craniala; Vertebrata; Euteleostomi;
OC Mammalia; Eutheria; Primates; Catarrhini; Homnidae; Homo.
OX NCBI_TaxID=9606;
RN (1)
RP SEQUENCE FROM N.A.
RC TISSUE=LIVER;
RA Wydro R., Cohen E., Dackowski W., Stenflo J., Lundwall A.,
RA Dahlback B.;
RL Submitted (FEB-1992) to the EMBL/GenBank/DBJ databases.
DR EMBL: X12892; CAJ31383.1; -
DR HSP: P00740; ICFH.
DR InterPro: IPR000152; ASX_hydroxyl.
DR InterPro: IPR000561; EGF-like.
DR InterPro: IPR001881; EGF_CA.
DR InterPro: IPR002383; GLA_blood.
DR InterPro: IPR001791; Laminin_G.
DR InterPro: IPR000294; VitK_dep_GLA.
DR Pfam: PF00008; EGF_2.
DR Pfam: PF00594; g1a; 1.
DR Pfam: PF00054; Laminin_G; 1.
DR PRINTS: PR00001; GLABLOOD.
DR SMART: SM00179; EGF_CA; 3.
DR SMART: SM00069; GLA; 1.
DR SMART: SM00282; LamG; 2.
DR PROSITE: PS00010; ASX_HYDROXYL; 3.
DR PROSITE: PS00022; EGF_1; UNKNOWN_1.
DR PROSITE: PS01186; EGF_2; 3.
DR PROSITE: PS01187; EGF_CA; 2.
DR PROSITE: PS00011; GLU_CARBOXYLATION; 1.
DR Calcium-binding; EGF-like domain; Glycoprotein; Hydroxylation; Repeat;
KW Signal.
FT SIGNAL 1 15 POTENTIAL.
FT CHAIN 16 650 POTENTIAL.
FT SEQUENCE 650 AA; 72480 MW; C67343ECEB645174 CRC64;

Query Match          43.1%; Score 85; DB 4; Length 650;
Best Local Similarity 38.6%; Pred. No. 6.1e-06;
Matches 17; Conservative 10; Mismatches 17; Indels 0; Gaps 0;

OY 1 ANSFLXLRGSLXRXIXICDFXAKXIFEDVDTLAWFSKH 44
Db 16 ANSLFEELKGNLERECNEETCSYEAREVVEDDTKNEFWN 82

RESULT 11
O16519 PRELIMINARY; PRT; 650 AA.
AC O16519;
DT 01-NOV-1996 (Tremblrel. 01, Created)

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DR 01-NOV-1996 (TREMblrel. 01, Last sequence update)
DE 01-DEC-2001 (TREMblrel. 19, Last annotation update)
DE Protein S precursor (Fragment).
GN PROS1.
OS Homo sapiens (Human).
OC Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC Mammalia; Eutheria; Primates; Catarrhini; Hominiidae; Homo.
OX NCBI_TaxID=9606;
RN [1]
RP SEQUENCE FROM N.A.
RX MEDLINE=86313649; PubMed=2944113;
RA Lundwall A., Dackowski W., Cohen E., Shaffer M., Mahr A., Dahlback B.,
RA Stenflo J., Wydro R.,
RT "Isolation and sequence of the cDNA for human protein S, a regulator
RT of blood coagulation."
RL Proc. Natl. Acad. Sci. U.S.A. 83:6716-6720(1986).
DR EMBL: M1438; AAA60181.1; -.
DR HSSP: P00740; ICFH.
DR InterPro: IPR000152; ASX_hydroxyl.
DR InterPro: IPR000561; EGF-like.
DR InterPro: IPR001881; EGF-Ca.
DR InterPro: IPR002383; GLA_blood.
DR InterPro: IPR001791; Laminin_G.
DR Pfam: PF00008; EGF; 4.
DR Pfam: PF00594; gla; 1.
DR Pfam: PF00054; Laminin_G; 1.
DR PRINTS: PR00001; GLABLOOD.
DR SMART: SM00179; EGF_CA; 3.
DR SMART: SM00069; GLA; 1.
DR SMART: SM00282; LamG; 2.
DR PROSITE: PS00010; ASX_HYDROXYL; 3.
DR PROSITE: PS00022; EGF_1; UNKNOWN_1.
DR PROSITE: PS01186; EGF_2; 3.
DR PROSITE: PS01187; EGF_CA; 2.
DR PROSITE: PS00011; GLU CARBOXYLATION; 1.
KW Calcium-binding; EGF-like domain; Glycoprotein; Hydroxylation; Repeat;
KW Signal.
FT NON_TER
FT SIGNAL
FT CHAIN
SQ SEQUENCE 650 AA; 72462 MW; 9A8C044C503BF474 CRC64;

Query Match
Best Local Similarity 43.1%; Score 85; DB 4; Length 650;
Matches 17; Conservative 10; Mismatches 17; Indels 0; Gaps 0;

OY 1 ANSFLXLRGSLXRCIXICDFXAKXIFEDVDTLAFWSKH 44
DB 16 ANSLLEETKGNLERECIEELCNKEAREVFENDPETDYFYFKY 59

RESULT 12
O15253 PRELIMINARY; PRT; 100 AA.
AC O15253;
DT 01-NOV-1996 (TREMblrel. 01, Created)
DT 01-NOV-1996 (TREMblrel. 01, Last sequence update)
DT 01-DEC-2001 (TREMblrel. 19, Last annotation update)
DE Thrombin precursor (Fragment).
GN F2.
OS Homo sapiens (Human).
OC Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC Mammalia; Eutheria; Primates; Catarrhini; Hominiidae; Homo.
OX NCBI_TaxID=9606;
RN [1]
RP SEQUENCE FROM N.A.
RX MEDLINE=87182874; PubMed=3471151.
RA MacGillivray R.T., Irwin D.M., Guinto E.R., Stone J.C.;
RT "Recombinant genetic approaches to functional mapping of thrombin.";
RL Ann. N.Y. Acad. Sci. 485:73-79(1986).
DR EMBL: M33031; AAA60220.1; -.
DR HSSP: P00735; 2PFI.

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DR InterPro: IPR002383; GLA_blood.
DR InterPro: IPR000294; VitK_dep_GLA.
DR Pfam: PF00594; gla; 1.
DR PRINTS: PR00001; GLABLOOD.
DR SMART: SM00069; GLA; 1.
DR PROSITE: PS00011; GLU_CARBOXYLATION; 1.
FT SIGNAL
FT CHAIN
FT NON_TER
SQ SEQUENCE 100 AA; 11302 MW; FDOE5D0174E1F6FE CRC64;

Query Match
Best Local Similarity 42.6%; Score 84; DB 4; Length 100;
Matches 16; Conservative 8; Mismatches 20; Indels 0; Gaps 0;

OY 1 ANSFLXLRGSLXRCIXICDFXAKXIFEDVDTLAFWSKH 44
DB 44 ANTFLEVAKGNLERECVETCSYEDAEALRSTATDVFWAKY 87

RESULT 13
O8T6I3 PRELIMINARY; PRT; 542 AA.
AC O8T6I3;
DT 01-JUN-2002 (TREMblrel. 21, Created)
DT 01-JUN-2002 (TREMblrel. 21, Last sequence update)
DT 01-JUN-2002 (TREMblrel. 21, Last annotation update)
DE Gla-like protein.
OC Halocynthia roretzi (Sea squirt).
OC Eukaryota; Metazoa; Chordata; Urochordata; Ascidiacea;
OC Stolidobranchia; Pyrosidae; Halocynthia.
OX NCBI_TaxID=7729;
RN [1]
RP SEQUENCE FROM N.A.
RX Wang C.-P., Stafford D.W.;
RT "Halocynthia roretzi gla-like protein partial genomic DNA sequence.";
RL Submitted (Apr-2002) to the EMBL/Genbank/DBJ databases.
DR EMBL: AF466701; AAL74247.2; -.
SQ SEQUENCE 542 AA; 62090 MW; EB9BF13FE42B32FE CRC64;

Query Match
Best Local Similarity 41.9%; Score 82.5; DB 5; Length 542;
Matches 15; Conservative 10; Mismatches 17; Indels 1; Gaps 1;

OY 3 SFLXLRGSLXRCIXICDFXAKXIFE-DVDDTLAFWSKH 44
DB 33 SHFEELQGNLERECYEEELCSFEAREVFETNIQDLNFWAKY 75

RESULT 14
O61109 PRELIMINARY; PRT; 446 AA.
AC O61109;
DT 01-NOV-1996 (TREMblrel. 01, Created)
DT 01-NOV-1996 (TREMblrel. 01, Last sequence update)
DT 01-JUN-2002 (TREMblrel. 21, Last annotation update)
DE Coagulation factor VII.
GN F7 OR FVII.
OS Mus musculus (Mouse).
OC Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC Mammalia; Eutheria; Rodentia; Sciurognathi; Muridae; Murinae; Mus.
OX NCBI_TaxID=10090;
RN [1]
RP SEQUENCE FROM N.A.
RX TISSUE=LIVER;
RX MEDLINE=96276538; PubMed=8701412;
RX Idusogie E., Rosen E., Geng J.P., Carmeliet P., Collen D.,
RA Castellino F.J.;
RT "Characterization of a cDNA encoding murine coagulation factor VII.";
RL Thromb. Haemost. 75:481-487(1996).
CC -1- SIMILARITY: BELONGS TO PEPTIDASE FAMILY S1; ALSO KNOWN AS THE
CC TRYPSIN FAMILY.

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DR EMBL: U44795; AAC52570.1; -  
 DR HSSP: P08709; 1FAK.  
 DR MEROPS: S01.215; -  
 DR MGD: MGI:109325; F7.  
 DR InterPro: IPR002086; Aldehyde\_dehydr.  
 DR InterPro: IPR000152; Asx\_hydroxyl.  
 DR InterPro: IPR001314; Chymotrypsin.  
 DR InterPro: IPR001064; Crystallin.  
 DR InterPro: IPR000561; EGF-like.  
 DR InterPro: IPR001881; EGF\_Ca.  
 DR InterPro: IPR002383; GLA\_blood.  
 DR InterPro: IPR001254; Ser\_protease\_Try.  
 DR InterPro: IPR000294; Vitk\_dep\_GLA.  
 DR Pfam: PF00008; EGF\_2.  
 DR Pfam: PF00594; gla; 1.  
 DR Pfam: PF00089; trypsin; 1.  
 DR PRINTS: PRO0722; CHYMOTRYPSIN.  
 DR PRINTS: PRO0001; GLABLOOD.  
 DR SMART: SM00179; EGF\_CA; 1.  
 DR SMART: SM00001; EGF\_like; 1.  
 DR SMART: SM00069; gla; 1.  
 DR SMART: SM00020; TRYP\_SPC; 1.  
 DR PROSITE: PS00070; ALDEHYDE\_DEHYDR\_CYS; UNKNOWN\_1.  
 DR PROSITE: PS00010; ASX\_HYDROXYL; UNKNOWN\_1.  
 DR PROSITE: PS00225; CRYSTALLIN\_BETAGAMMA; UNKNOWN\_1.  
 DR PROSITE: PS00022; EGF\_1; UNKNOWN\_1.  
 DR PROSITE: PS01187; EGF\_CA; 1.  
 DR PROSITE: PS00011; GLU\_CARBOXYLATION; 1.  
 DR PROSITE: PS50240; TRYPsin\_DOM; 1.  
 DR PROSITE: PS00134; TRYPsin\_HIS; UNKNOWN\_1.  
 DR PROSITE: PS00135; TRYPsin\_SER; 1.  
 DR Calcium-binding; EGF-like domain; Glycoprotein; Hydrolase; Repeat;  
 KW Serine protease.  
 SO SEQUENCE 446 AA; 50318 MW; 482FD09BEFDA6870 CRC64;

Query Match 40.6%; Score 80; DB 11; Length 446;  
 Best Local Similarity 43.9%; Pred. No. 2.9e-05;  
 Matches 18; Conservative 3; Mismatches 20; Indels 0; Gaps 0;

QY 1 ANSFLXLRGSLRKXIXXICDFXXAKXIFEDVDTLAW 41  
 DB 42 ANSLLELPWGLERECNEQCSFEARELFKSPERTKQW 82

## RESULT 15

ID Q14316 PRELIMINARY; PRT; 456 AA;  
 AC Q14316;  
 DT 01-NOV-1996 (Tremblrel. 01, Created)  
 DT 01-AUG-1999 (Tremblrel. 11, Last sequence update)  
 DT 01-JUN-2002 (Tremblrel. 21, Last annotation update)  
 DE P9 (coagulation factor IX (Plasma THROMBOPLASTIC component, Christmas disease, HAEMOPHILIA B)) (Factor IX).  
 GN F9 OR FACTOR IX.  
 GN F9 OR FACTOR IX.  
 OS Homo sapiens (Human).  
 OC Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
 OC Mammalia; Eutheria; Primates; Catarrhini; Homnidae; Homo.  
 OC NCBI\_TaxID=9606;  
 RN (1)  
 RP SEQUENCE FROM N.A.  
 RA Bird C.;  
 RL Submitted (NOV-1998) to the EMBL/GenBank/DBJ databases.  
 RN [2]  
 RP SEQUENCE OF 3-19 FROM N.A.  
 RX MEDLINE=88377116; PubMed=3416069;  
 RA Reitema P.A., Bertina R.M., Ploos van Amstel J.K., Riemens A.,  
 RA Briel E.;  
 RT "The putative factor IX gene promoter in hemophilia B Leyden";  
 RL Blood 72:1074-1076(1988)  
 CC -1- SIMILARTY: BELONGS TO PEPTIDASE FAMILY S1; ALSO KNOWN AS THE  
 CC TRYPSIN FAMILY.  
 DR EMBL: AL033403; CAA21954.1; -  
 DR EMBL: X55008; CAB38245.2; -

DR HSSP: P00740; 1CFH.  
 DR MEROPS: S01.214; -  
 DR InterPro: IPR000152; Asx\_hydroxyl.  
 DR InterPro: IPR001314; Chymotrypsin.  
 DR InterPro: IPR000561; EGF-like.  
 DR InterPro: IPR000742; EGF\_2.  
 DR InterPro: IPR001881; EGF\_Ca.  
 DR InterPro: IPR001438; EGF\_II.  
 DR InterPro: IPR002383; GLA\_blood.  
 DR InterPro: IPR001254; Ser\_protease\_Try.  
 DR InterPro: IPR000294; Vitk\_dep\_GLA.  
 DR Pfam: PF00594; gla; 1.  
 DR Pfam: PF00089; trypsin; 1.  
 DR PRINTS: PRO0722; CHYMOTRYPSIN.  
 DR PRINTS: PRO0001; GLABLOOD.  
 DR SMART: SM00179; EGF\_CA; 1.  
 DR SMART: SM00001; EGF\_like; 1.  
 DR SMART: SM00069; gla; 1.  
 DR SMART: SM00020; TRYP\_SPC; 1.  
 DR PROSITE: PS00010; ASX\_HYDROXYL; UNKNOWN\_1.  
 DR PROSITE: PS00022; EGF\_1; UNKNOWN\_1.  
 DR PROSITE: PS01186; EGF\_2; 2.  
 DR PROSITE: PS01187; EGF\_CA; 1.  
 DR PROSITE: PS00011; GLU\_CARBOXYLATION; 1.  
 DR PROSITE: PS50240; TRYPsin\_DOM; 1.  
 DR PROSITE: PS00134; TRYPsin\_HIS; UNKNOWN\_1.  
 DR PROSITE: PS00135; TRYPsin\_SER; 1.  
 DR Calcium-binding; EGF-like domain; Glycoprotein; Hydrolase; Repeat;  
 KW Serine protease.  
 SO SEQUENCE 456 AA; 51149 MW; 54E20A1B3964E234 CRC64;

Query Match 40.6%; Score 80; DB 4; Length 456;  
 Best Local Similarity 37.1%; Pred. No. 3e-05;  
 Matches 13; Conservative 8; Mismatches 14; Indels 0; Gaps 0;

QY 10 QGSLRXKIXXICDFXXAKXIFEDVDTLAFWSKH 44  
 DB 52 QGNLERCEMEKCSFEAREVFEVTERTEWKKY 86

Search completed: May 16, 2003, 10:15:27  
 Job time : 31 secs

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GenCore version 5.1.4-p5-4578  
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OM protein - protein search, using sw model

Run on: May 16, 2003, 10:14:04 ; Search time 14 Seconds  
(without alignments)  
92.472 Million cell updates/sec

Title: SEQ1-4EDITS  
Perfect score: 197  
Sequence: 1 ANSEFLXLRGSLRXKXICIX.....XXAKXIFEDVDTLAFWSKH 44

Scoring table: BLOSUM62  
Gapop 10.0 , Gapext 0.5

Searched: 262574 seqs, 29422922 residues

Total number of hits satisfying chosen parameters: 262574

Minimum DB seq length: 0  
Maximum DB seq length: 2000000000

Post-processing: Minimum Match 0%  
Maximum Match 100%  
Listing first 45 summaries

Database : Issued\_Patents\_AA:  
1: /cgn2\_6/ptodata/1/1aa/5A.COMB.pep:\*  
2: /cgn2\_6/ptodata/1/1aa/5B.COMB.pep:\*  
3: /cgn2\_6/ptodata/1/1aa/6A.COMB.pep:\*  
4: /cgn2\_6/ptodata/1/1aa/6B.COMB.pep:\*  
5: /cgn2\_6/ptodata/1/1aa/PTOTS.COMB.pep:\*  
6: /cgn2\_6/ptodata/1/1aa/Backfiles1.pep:\*

Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

## SUMMARIES

Result No.	Score	Query Match	Length	ID	Description
1	179	90.9	44	US-08-955-636-24	Sequence 24, Appl
2	176	89.3	44	US-08-955-636-35	Sequence 35, Appl
3	173	87.8	44	US-08-955-636-20	Sequence 20, Appl
4	170	86.3	44	US-08-955-636-21	Sequence 19, Appl
5	168	85.3	44	US-08-955-636-19	Sequence 19, Appl
6	168	85.3	44	US-08-955-636-22	Sequence 22, Appl
7	160	81.2	44	US-08-955-636-1	Sequence 1, Appl
8	160	81.2	44	US-08-955-636-25	Sequence 25, Appl
9	160	81.2	45	US-08-965-832-2	Sequence 2, Appl
10	160	81.2	419	US-08-295-411-1	Sequence 1, Appl
11	160	81.2	419	US-08-955-471-1	Sequence 3, Appl
12	160	81.2	419	US-09-667-570A-3	Sequence 1, Appl
13	160	81.2	419	PCT-US92-10242-1	Sequence 1, Appl
14	160	81.2	460	US-08-756-506-2	Sequence 4, Appl
15	160	81.2	460	US-08-756-506-4	Sequence 4, Appl
16	160	81.2	460	5270178-13	Patent No. 5270178
17	160	81.2	460	5270178-14	Patent No. 5270178
18	160	81.2	460	5270178-15	Patent No. 5270178
19	160	81.2	460	5270178-16	Patent No. 5270178
20	160	81.2	461	5225537-2	Patent No. 5225537
21	160	81.2	461	5270178-17	Patent No. 5270178
22	160	81.2	461	5270178-18	Patent No. 5270178
23	160	81.2	461	5460953-3	Patent No. 5460953
24	147	74.6	44	US-08-745-254A-2	Sequence 2, Appl
25	147	74.6	461	5270178-2	Patent No. 5270178
26	143	72.6	41	US-08-229-280-5	Sequence 5, Appl
27	129	65.5	409	US-09-065-872-2	Sequence 2, Appl

28	129	65.5	409	4	US-09-667-570A-2	Sequence 2, Appl
29	129	65.5	410	4	US-09-065-872-1	Sequence 1, Appl
30	129	65.5	410	4	US-09-667-570A-1	Sequence 1, Appl
31	117	59.4	44	3	US-08-955-636-23	Sequence 23, Appl
32	116	58.9	44	3	US-08-955-636-2	Sequence 2, Appl
33	114	57.9	139	1	US-08-330-978-2	Sequence 2, Appl
34	114	57.9	139	1	US-08-474-042-2	Sequence 2, Appl
35	114	57.9	139	1	US-08-484-558-2	Sequence 2, Appl
36	114	57.9	139	1	US-08-774-592-2	Sequence 2, Appl
37	114	57.9	437	1	US-08-487-037-2	Sequence 2, Appl
38	114	57.9	437	1	US-08-487-037-3	Sequence 3, Appl
39	114	57.9	487	1	US-08-469-486-53	Sequence 53, Appl
40	114	57.9	487	2	US-08-469-658-53	Sequence 53, Appl
41	114	57.9	488	1	US-08-487-037-1	Sequence 1, Appl
42	114	57.9	492	1	US-08-469-486-2	Sequence 2, Appl
43	114	57.9	492	2	US-08-469-658-2	Sequence 2, Appl
44	110	55.8	448	1	US-08-295-411-3	Sequence 3, Appl
45	110	55.8	448	2	US-08-955-471-3	Sequence 3, Appl

## ALIGNMENTS

```

RESULT 1
US-08-955-636-24
Sequence 24, Application US/08955636A
Patent No. 6017882
GENERAL INFORMATION:
APPLICANT: Neilsen, Gary
TITLE OF INVENTION: MODIFIED VITAMIN K-DEPENDENT
FILE REFERENCE: 09531/002001
CURRENT APPLICATION NUMBER: US/08/955,636A
CURRENT FILING DATE: 1997-10-23
NUMBER OF SEQ ID NOS: 35
SOFTWARE: FastSeq for Windows Version 3.0
SEQ ID NO 24
LENGTH: 44
TYPE: PRT
ORGANISM: Homo sapiens
FEATURE:
NAME/KEY: MOD_RES
LOCATION: (0)...(0)
OTHER INFORMATION: Xaa-gamma carboxyglutamic acid or glutamic acid
US-08-955-636-24

Query Match          90.9%; Score 179; DB 3; Length 44;
Best Local Similarity 100.0%; Pred. No. 1.2e-23;
Matches 44; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY      1 ANSEFLXLRGSLRXKXICIXCDPFXAKXIFEDVDTLAFWSKH 44
DB      1 ANSEFLXLRGSLRXKXICIXCDPFXAKXIFEDVDTLAFWSKH 44

RESULT 2
US-08-955-636-35
Sequence 35, Application US/08955636A
Patent No. 6017882
GENERAL INFORMATION:
APPLICANT: Neilsen, Gary
TITLE OF INVENTION: MODIFIED VITAMIN K-DEPENDENT
FILE REFERENCE: 09531/002001
CURRENT APPLICATION NUMBER: US/08/955,636A
CURRENT FILING DATE: 1997-10-23
SOFTWARE: FastSeq for Windows Version 3.0
SEQ ID NO 35
LENGTH: 44
TYPE: PRT
ORGANISM: Homo sapiens
FEATURE:

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NAME/KEY: MOD\_RES  
LOCATION: (0)...(0)  
OTHER INFORMATION: xaa-gamma carboxyglutamic acid or glutamic acid  
US-08-955-636-35

Query Match 89.3%; Score 176; DB 3; Length 44;  
Best Local Similarity 97.7%; Pred. No. 3.8e-23;  
Matches 43; Conservative 1; Mismatches 0; Indels 0; Gaps 0;

QY 1 ANSFLXLRQSLKRXKXCIXXICDFXXAKXIFEDVDTLAFWSKH 44  
Db 1 ANSFLXLRQSLKRXKXCIXXICDFXXAKXIFEDVDTLAFWSKH 44

RESULT 3  
US-08-955-636-20  
Sequence 20, Application US/08955636A  
Patent No. 6017882  
GENERAL INFORMATION:  
APPLICANT: Nelstuen, Gary  
TITLE OF INVENTION: MODIFIED VITAMIN K-DEPENDENT  
FILE REFERENCE: 09531/002001  
CURRENT APPLICATION NUMBER: US/08/955,636A  
CURRENT FILING DATE: 1997-10-23  
NUMBER OF SEQ ID NOS: 35  
SOFTWARE: FastSeq for Windows Version 3.0  
SEQ ID NO 20  
LENGTH: 44  
TYPE: PRT  
ORGANISM: Homo sapiens  
FEATURE:  
NAME/KEY: MOD\_RES  
LOCATION: (0)...(0)  
OTHER INFORMATION: xaa-gamma carboxyglutamic acid or glutamic acid  
US-08-955-636-20

Query Match 87.8%; Score 173; DB 3; Length 44;  
Best Local Similarity 97.7%; Pred. No. 1.2e-22;  
Matches 43; Conservative 0; Mismatches 1; Indels 0; Gaps 0;

QY 1 ANSFLXLRQSLKRXKXCIXXICDFXXAKXIFEDVDTLAFWSKH 44  
Db 1 ANSFLXLRQSLKRXKXCIXXICDFXXAKXIFEDVDTLAFWSKH 44

RESULT 4  
US-08-955-636-21  
Sequence 21, Application US/08955636A  
Patent No. 6017882  
GENERAL INFORMATION:  
APPLICANT: Nelstuen, Gary  
TITLE OF INVENTION: MODIFIED VITAMIN K-DEPENDENT  
FILE REFERENCE: 09531/002001  
CURRENT APPLICATION NUMBER: US/08/955,636A  
CURRENT FILING DATE: 1997-10-23  
NUMBER OF SEQ ID NOS: 35  
SOFTWARE: FastSeq for Windows Version 3.0  
SEQ ID NO 21  
LENGTH: 44  
TYPE: PRT  
ORGANISM: Homo sapiens  
FEATURE:  
NAME/KEY: MOD\_RES  
LOCATION: (0)...(0)  
OTHER INFORMATION: xaa-gamma carboxyglutamic acid or glutamic acid  
US-08-955-636-21

Query Match 86.3%; Score 170; DB 3; Length 44;  
Best Local Similarity 95.5%; Pred. No. 4e-22;  
Matches 42; Conservative 1; Mismatches 1; Indels 0; Gaps 0;

QY 1 ANSFLXLRQSLKRXKXCIXXICDFXXAKXIFEDVDTLAFWSKH 44  
Db 1 ANSFLXLRQSLKRXKXCIXXICDFXXAKXIFEDVDTLAFWSKH 44

RESULT 5  
US-08-955-636-19  
Sequence 19, Application US/08955636A  
Patent No. 6017882  
GENERAL INFORMATION:  
APPLICANT: Nelstuen, Gary  
TITLE OF INVENTION: MODIFIED VITAMIN K-DEPENDENT  
FILE REFERENCE: 09531/002001  
CURRENT APPLICATION NUMBER: US/08/955,636A  
CURRENT FILING DATE: 1997-10-23  
NUMBER OF SEQ ID NOS: 35  
SOFTWARE: FastSeq for Windows Version 3.0  
SEQ ID NO 19  
LENGTH: 44  
TYPE: PRT  
ORGANISM: Homo sapiens  
FEATURE:  
NAME/KEY: MOD\_RES  
LOCATION: (0)...(0)  
OTHER INFORMATION: xaa-gamma carboxyglutamic acid or glutamic acid  
US-08-955-636-19

Query Match 85.3%; Score 168; DB 3; Length 44;  
Best Local Similarity 95.5%; Pred. No. 8.9e-22;  
Matches 42; Conservative 0; Mismatches 2; Indels 0; Gaps 0;

QY 1 ANSFLXLRQSLKRXKXCIXXICDFXXAKXIFEDVDTLAFWSKH 44  
Db 1 ANSFLXLRQSLKRXKXCIXXICDFXXAKXIFEDVDTLAFWSKH 44

RESULT 6  
US-08-955-636-22  
Sequence 22, Application US/08955636A  
Patent No. 6017882  
GENERAL INFORMATION:  
APPLICANT: Nelstuen, Gary  
TITLE OF INVENTION: MODIFIED VITAMIN K-DEPENDENT  
FILE REFERENCE: 09531/002001  
CURRENT APPLICATION NUMBER: US/08/955,636A  
CURRENT FILING DATE: 1997-10-23  
NUMBER OF SEQ ID NOS: 35  
SOFTWARE: FastSeq for Windows Version 3.0  
SEQ ID NO 22  
LENGTH: 44  
TYPE: PRT  
ORGANISM: Homo sapiens  
FEATURE:  
NAME/KEY: MOD\_RES  
LOCATION: (0)...(0)  
OTHER INFORMATION: xaa-gamma carboxyglutamic acid or glutamic acid  
US-08-955-636-22

Query Match 85.3%; Score 168; DB 3; Length 44;  
Best Local Similarity 95.5%; Pred. No. 8.9e-22;  
Matches 42; Conservative 0; Mismatches 2; Indels 0; Gaps 0;

QY 1 ANSFLXLRQSLKRXKXCIXXICDFXXAKXIFEDVDTLAFWSKH 44  
Db 1 ANSFLXLRQSLKRXKXCIXXICDFXXAKXIFEDVDTLAFWSKH 44

RESULT 7  
US-08-955-636-1  
Sequence 1, Application US/08955636A  
Patent No. 6017882

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: GENERAL INFORMATION:
: APPLICANT: Nelsestuen, Gary
: TITLE OF INVENTION: MODIFIED VITAMIN K-DEPENDENT
: FILE REFERENCE: 09531/002001
: CURRENT APPLICATION NUMBER: US/08/955,636A
: CURRENT FILING DATE: 1997-10-23
: NUMBER OF SEQ ID NOS: 35
: SOFTWARE: FASTSEQ for Windows Version 3.0
: SEQ ID NO: 1
: TYPE: PRP
: ORGANISM: Homo sapiens
: FEATURE:
: NAME/KEY: MOD_RES
: LOCATION: (0)...(0)
: OTHER INFORMATION: Xaa-gamma carboxyglutamic acid or glutamic acid
US-08-955-636-1

Query Match
Best Local Similarity 81.2%; Score 160; DB 3; Length 44;
Matches 40; Conservative 2; Mismatches 2; Indels 0; Gaps 0;

OY 1 ANSFLXXLRQSLXRCIXXICDFXXAKXIFEDVDDTLAFWSKH 44
DB 1 ANSFLXXLRHSSLXRCIXXICDFXXAKXIFQVNDTLAFWSKH 44

RESULT 8
US-08-955-636-25
: Sequence 25, Application US/08955636A
: Patent No. 6017882
: GENERAL INFORMATION:
: APPLICANT: Nelsestuen, Gary
: TITLE OF INVENTION: MODIFIED VITAMIN K-DEPENDENT
: FILE REFERENCE: 09531/002001
: CURRENT APPLICATION NUMBER: US/08/955,636A
: CURRENT FILING DATE: 1997-10-23
: NUMBER OF SEQ ID NOS: 35
: SOFTWARE: FASTSEQ for Windows Version 3.0
: SEQ ID NO: 25
: TYPE: PRP
: ORGANISM: Homo sapiens
: FEATURE:
: NAME/KEY: MOD_RES
: LOCATION: (0)...(0)
: OTHER INFORMATION: Xaa-gamma carboxyglutamic acid or glutamic acid
US-08-955-636-25

Query Match
Best Local Similarity 93.2%; Score 160; DB 3; Length 44;
Matches 41; Conservative 0; Mismatches 3; Indels 0; Gaps 0;

OY 1 ANSFLXXLRQSLXRCIXXICDFXXAKXIFEDVDDTLAFWSKH 44
DB 1 ANSFLXXLRHSSLXRCIXXICDFXXAKXIFEDVDDTLAFWSKH 44

RESULT 9
US-08-965-832-2
: Sequence 2, Application US/08965832
: Patent No. 5847085
: GENERAL INFORMATION:
: APPLICANT: CHARLES T. ESMON AND MIKHAIL D. SMIRNOV
: TITLE OF INVENTION: Modified Protein C
: NUMBER OF SEQUENCES: 4
: CORRESPONDENCE ADDRESS:
: ADDRESSEE: Patrea L. Pabst
: STREET: 2800 One Atlantic Center, 1201 West
: STREET: Peachtree Street
: CITY: Atlanta
```

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: STATE: GA
: COUNTRY: USA
: ZIP: 30309-3450
: COMPUTER READABLE FORM:
: MEDIUM TYPE: Floppy disk
: COMPUTER: IBM PC compatible
: OPERATING SYSTEM: PC-DOS/MS-DOS
: SOFTWARE: PatentIn Release #1.0, Version #1.25
: CURRENT APPLICATION DATA:
: APPLICATION NUMBER: US/08/965,832
: FILING DATE: 7-NOV-1997
: CLASSIFICATION: 530
: PRIOR APPLICATION DATA:
: APPLICATION NUMBER: 08/745,254
: FILING DATE: 8-NOV-1996
: PRIOR APPLICATION DATA:
: APPLICATION NUMBER: 60/053,768
: FILING DATE: 25-JUL-1997
: ATTORNEY/AGENT INFORMATION:
: NAME: Pabst, Patrea L.
: REGISTRATION NUMBER: 31,284
: REFERENCE/DOCKET INFORMATION:
: TELECOMMUNICATION INFORMATION:
: TELEPHONE: (404)-873-8794
: TELEFAX: (404)-873-8795
: INFORMATION FOR SEQ ID NO: 2:
: SEQUENCE CHARACTERISTICS:
: LENGTH: 45 amino acids
: TYPE: amino acid
: STRANDEDNESS: single
: TOPOLOGY: linear
: MOLECULE TYPE: Peptide
: FEATURE:
: NAME/KEY:
: LOCATION: 6, 7, 14, 16, 19, 20, 25, 26, 29
: OTHER INFORMATION: /note="where Xaa means gamma
: FEATURE:
: NAME/KEY:
: LOCATION:
: OTHER INFORMATION: /note="partial sequence of human protein C"
US-08-965-832-2

Query Match
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Matches 40; Conservative 2; Mismatches 2; Indels 0; Gaps 0;

OY 1 ANSFLXXLRQSLXRCIXXICDFXXAKXIFEDVDDTLAFWSKH 44
DB 1 ANSFLXXLRHSSLXRCIXXICDFXXAKXIFQVNDTLAFWSKH 44

RESULT 10
US-08-295-411-1
: Sequence 1, Application US/08295411
: Patent No. 5679639
: GENERAL INFORMATION:
: APPLICANT: Griffith, John H.
: APPLICANT: Mesters, Rolf M.
: TITLE OF INVENTION: Serine Protease-Derived Polypeptides and
: TITLE OF INVENTION: Anti-Peptide Antibodies, Systems and Therapeutic Methods
: TITLE OF INVENTION: for Inhibiting Coagulation
: NUMBER OF SEQUENCES: 10
: CORRESPONDENCE ADDRESS:
: ADDRESSEE: Office of Patent Counsel, The Scripps
: STREET: Research Institute
: STREET: 10666 No. 5679639th Torrey Pines Road, TPC 8
: CITY: La Jolla
: STATE: CA
: COUNTRY: USA
: ZIP: 92037
: COMPUTER READABLE FORM:
: MEDIUM TYPE: Floppy disk
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COMPUTER: IBM PC compatible  
OPERATING SYSTEM: PC-DOS/MS-DOS  
SOFTWARE: Patentin Release #1.0, Version #1.25  
CURRENT APPLICATION DATA:  
FILING DATE: 18-NOV-1991  
APPLICATION NUMBER: US/08/295,411  
FILING DATE: 22-AUG-1994  
CLASSIFICATION: 530  
PRIOR APPLICATION DATA:  
APPLICATION NUMBER: US 07/793,989  
FILING DATE: 18-NOV-1991  
CLASSIFICATION: 530  
ATTORNEY/AGENT INFORMATION:  
NAME: Fitting, Thomas  
REGISTRATION NUMBER: 34,163  
REFERENCE/DOCKET NUMBER: TSRI263.0C1  
TELECOMMUNICATION INFORMATION:  
TELEPHONE: 619-554-2937  
TELEFAX: 619-554-6312  
INFORMATION FOR SEQ ID NO: 1:  
SEQUENCE CHARACTERISTICS:  
LENGTH: 419 amino acids  
TYPE: amino acid  
TOPOLOGY: linear  
MOLECULE TYPE: protein  
HYPOTHETICAL: NO  
ANTI-SENSE: NO  
FEATURE:  
NAME/KEY: Region  
LOCATION: 1..157  
OTHER INFORMATION: /note= "Protein C Light Chain"  
FEATURE:  
NAME/KEY: Region  
LOCATION: 158..169  
OTHER INFORMATION: /note= "Protein C Activation"  
OTHER INFORMATION: Peptide"  
FEATURE:  
NAME/KEY: Region  
LOCATION: 170..419  
OTHER INFORMATION: /note= "Protein C Heavy Chain"  
US-08-295-411-1

Query Match 81.2%; Score 160; DB 1; Length 419;  
Best Local Similarity 70.5%; Pred. No. 2.7e-19;  
Matches 31; Conservative 2; Mismatches 11; Indels 0; Gaps 0;

Qy 1 ANSFLXLRQGSIXKXICDFXKXKXIFEDVDDTLAEWSKH 44  
Db 1 ANSFLERHSSLERECIEIEEAKKEIFQNVDDTLAEWSKH 44

RESULT 11  
US-08-955-471-1  
Sequence 1, Application US/08955471  
Patent No. 5968751  
GENERAL INFORMATION:  
APPLICANT: Gritfin, John H.  
APPLICANT: Mesters, Rolf M.  
TITLE OF INVENTION: Serine Protease-Derived Polypeptides and  
TITLE OF INVENTION: Anti-Peptide Antibodies, Systems and Therapeutic Methods  
TITLE OF INVENTION: for Inhibiting Coagulation  
NUMBER OF SEQUENCES: 10  
CORRESPONDENCE ADDRESS:  
ADDRESSEE: Office of Patent Counsel, The Scripps  
ADDRESSEE: Research Institute  
STREET: 10666 No. 5968751th Torrey Pines Road, TPC 8  
CITY: La Jolla  
STATE: CA  
COUNTRY: USA  
ZIP: 92037  
COMPUTER READABLE FORM:  
MEDIUM TYPE: Floppy disk  
COMPUTER: IBM PC compatible  
OPERATING SYSTEM: PC-DOS/MS-DOS

SOFTWARE: Patentin Release #1.0, Version #1.25  
CURRENT APPLICATION DATA:  
APPLICATION NUMBER: US/08/955,471  
FILING DATE:  
CLASSIFICATION:  
PRIOR APPLICATION DATA:  
APPLICATION NUMBER: 08/295,411  
FILING DATE:  
CLASSIFICATION:  
ATTORNEY/AGENT INFORMATION:  
NAME: Fitting, Thomas  
REGISTRATION NUMBER: 34,163  
REFERENCE/DOCKET NUMBER: TSRI263.0C1  
TELECOMMUNICATION INFORMATION:  
TELEPHONE: 619-554-2937  
TELEFAX: 619-554-6312  
INFORMATION FOR SEQ ID NO: 1:  
SEQUENCE CHARACTERISTICS:  
LENGTH: 419 amino acids  
TYPE: amino acid  
TOPOLOGY: linear  
MOLECULE TYPE: protein  
HYPOTHETICAL: NO  
ANTI-SENSE: NO  
FEATURE:  
NAME/KEY: Region  
LOCATION: 1..157  
OTHER INFORMATION: /note= "Protein C Light Chain"  
FEATURE:  
NAME/KEY: Region  
LOCATION: 158..169  
OTHER INFORMATION: /note= "Protein C Activation"  
OTHER INFORMATION: Peptide"  
FEATURE:  
NAME/KEY: Region  
LOCATION: 170..419  
OTHER INFORMATION: /note= "Protein C Heavy Chain"  
US-08-955-471-1

Query Match 81.2%; Score 160; DB 2; Length 419;  
Best Local Similarity 70.5%; Pred. No. 2.7e-19;  
Matches 31; Conservative 2; Mismatches 11; Indels 0; Gaps 0;

Qy 1 ANSFLXLRQGSIXKXICDFXKXKXIFEDVDDTLAEWSKH 44  
Db 1 ANSFLERHSSLERECIEIEEAKKEIFQNVDDTLAEWSKH 44

RESULT 12  
US-09-667-570A-3  
Sequence 3, Application US/09667570A  
Patent No. 6436397  
GENERAL INFORMATION:  
APPLICANT: Baker, Jeffrey C  
APPLICANT: Carlson, Andrew D  
APPLICANT: Huang, Lihua  
APPLICANT: Shelliga, Theodore A  
TITLE OF INVENTION: Improved Methods for Processing Activated Protein C  
FILE REFERENCE: X-11796A  
CURRENT APPLICATION NUMBER: US/09/667,570A  
CURRENT FILING DATE: 2000-09-21  
PRIOR APPLICATION NUMBER: 60/045,255  
PRIOR FILING DATE: 1997-04-28  
NUMBER OF SEQ ID NOS: 3  
SOFTWARE: Patentin version 3.1  
SEQ ID NO 3  
LENGTH: 419  
TYPE: PRT  
ORGANISM: Homo sapiens  
US-09-667-570A-3

Query Match 81.2%; Score 160; DB 4; Length 419;  
Best Local Similarity 70.5%; Pred. No. 2.7e-19;





Fri May 16 11:31:06 2003

seq1-4edits.ra

Page 6

STREET: 1201 Eastlake Avenue East  
CITY: Seattle  
STATE: WA  
COUNTRY: USA  
ZIP: 98102  
COMPUTER READABLE FORM:  
MEDIUM TYPE: Floppy disk  
COMPUTER: IBM PC compatible  
OPERATING SYSTEM: PC-DOS/MS-DOS  
SOFTWARE: Patent In Release #1.0, Version #1.25  
CURRENT APPLICATION DATA:  
APPLICATION NUMBER: US/08/756,506  
FILING DATE:  
CLASSIFICATION: 800  
ATTORNEY/AGENT INFORMATION:  
NAME: Sawislak, Deborah A  
REGISTRATION NUMBER: 37,438  
REFERENCE/DOCKET NUMBER: 95-28  
TELECOMMUNICATION INFORMATION:  
TELEPHONE: 206-442-6672  
TELEFAX: 206-442-6678  
INFORMATION FOR SEQ ID NO: 4:  
SEQUENCE CHARACTERISTICS:  
LENGTH: 460 amino acids  
type: amino acid  
TOPOLOGY: linear  
MOLECULE TYPE: protein  
US-08-756-506-4

GenCore version 5.1.4-5-4578  
Copyright (c) 1993 - 2003 Compugen Ltd.

OM protein - protein search, using sw model

Run on: May 16, 2003, 10:15:34 : Search time 55 Seconds  
(without alignments)  
77.161 Million cell updates/sec

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Perfect score: 197  
Sequence: 1 ANSFLXLRQGSIXRXCIXX.....XXAKXIFEDVDITLAFWSKH 44

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Gapop 10.0 , Gapext 0.5

Searched: 362588 seqs, 96450795 residues

Total number of hits satisfying chosen parameters: 362588

Minimum DB seq length: 0  
Maximum DB seq length: 200000000

Post-processing: Minimum Match 0%  
Maximum Match 100%  
Listing first 45 summaries

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2: /cgn2\_6/ptodata/1/pubpaa/PCT\_NEW\_PUB.pep.\*  
3: /cgn2\_6/ptodata/1/pubpaa/US06\_NEW\_PUB.pep.\*  
4: /cgn2\_6/ptodata/1/pubpaa/US07\_NEW\_PUB.pep.\*  
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10: /cgn2\_6/ptodata/1/pubpaa/US05\_PUBCOMB.pep.\*  
11: /cgn2\_6/ptodata/1/pubpaa/US10\_NEW\_PUB.pep.\*  
12: /cgn2\_6/ptodata/1/pubpaa/US10\_PUBCOMB.pep.\*  
13: /cgn2\_6/ptodata/1/pubpaa/US60\_NEW\_PUB.pep.\*  
14: /cgn2\_6/ptodata/1/pubpaa/US60\_PUBCOMB.pep.\*

Pred. No. is the number of results predicted by chance to have a  
score greater than or equal to the score of the result being printed,  
and is derived by analysis of the total score distribution.

## SUMMARIES

Result No.	Score	Query Match	Length	ID	Description
1	179	90.9	419	9	US-10-182-263-6
2	174	88.3	419	9	US-10-182-263-3
3	174	88.3	419	9	US-10-182-263-4
4	174	88.3	419	9	US-10-182-263-5
5	160	81.2	419	9	US-10-182-263-1
6	160	81.2	419	9	US-09-978-917A-4
7	160	81.2	461	9	US-10-182-263-2
8	160	81.2	461	9	US-09-978-917A-2
9	99	50.3	466	9	US-10-017-122-2
10	96	48.7	406	9	US-10-109-498-1
11	84.5	42.9	96	9	US-09-759-130B-313
12	84.5	42.9	96	9	US-10-189-123-43
13	84.5	42.9	209	9	US-09-759-130B-312
14	84.5	42.9	209	9	US-10-189-123-42
15	84.5	42.9	226	9	US-09-759-130B-310
16	84.5	42.9	226	9	US-10-189-123-40
17	80	40.6	415	10	US-09-118-748-2
18	80	40.6	461	9	US-10-132-829-5
19	80	40.6	461	10	US-09-884-901-3

20	64.5	32.7	95	9	US-09-759-130B-356	Sequence 356, App
21	64.5	32.7	95	9	US-10-189-123-86	Sequence 86, Appl
22	64.5	32.7	208	9	US-09-759-130B-355	Sequence 355, App
23	64.5	32.7	208	9	US-10-189-123-85	Sequence 85, Appl
24	64.5	32.7	225	9	US-09-759-130B-353	Sequence 353, App
25	64.5	32.7	225	9	US-10-189-123-83	Sequence 83, Appl
26	49	24.9	1363	9	US-09-375-248-19	Sequence 19, Appl
27	48	24.4	348	10	US-09-982-610-18	Sequence 18, Appl
28	48	24.4	1298	10	US-09-982-610-33	Sequence 33, Appl
29	48	24.4	1363	9	US-09-375-248-2	Sequence 2, Appl
30	47	23.9	180	10	US-09-766-678-6	Sequence 6, Appl
31	47	23.9	317	9	US-09-939-833-5	Sequence 5, Appl
32	47	23.9	317	10	US-09-939-754-5	Sequence 5, Appl
33	47	23.9	317	10	US-09-939-833-5	Sequence 5, Appl
34	47	23.9	367	9	US-09-939-833-12	Sequence 12, Appl
35	47	23.9	367	10	US-09-939-754-12	Sequence 12, Appl
36	47	23.9	367	10	US-09-939-833-12	Sequence 12, Appl
37	47	23.9	1356	9	US-09-969-037-7	Sequence 7, Appl
38	47	23.9	1356	9	US-10-022-939-2	Sequence 2, Appl
39	47	23.9	1356	9	US-10-100-405A-2	Sequence 2, Appl
40	47	23.9	1367	10	US-09-919-408-6	Sequence 6, Appl
41	47	23.9	1367	10	US-09-766-678-2	Sequence 2, Appl
42	47	23.9	1367	10	US-09-872-136-6	Sequence 6, Appl
43	45	22.8	254	10	US-09-796-149-4	Sequence 4, Appl
44	44.5	22.6	49	9	US-09-836-392-34	Sequence 34, Appl
45	44	22.3	52	9	US-09-796-692-2179	Sequence 2179, Ap

## ALIGNMENTS

RESULT 1  
US-10-182-263-6  
; Sequence 6, Application US/10182263  
; Publication No. US20030022354A1  
; GENERAL INFORMATION:  
; APPLICANT: Gerlitz, Bruce E  
; APPLICANT: Jones, Bryan E  
; APPLICANT: Grinnell, Brian W  
; TITLE OF INVENTION: PROTEIN C DERIVATIVES  
; FILE REFERENCE: X-13611  
; CURRENT APPLICATION NUMBER: US/10/182,263  
; CURRENT FILING DATE: 2002-07-22  
; PRIOR APPLICATION NUMBER: 60/181948  
; PRIOR FILING DATE: 2002-02-11  
; PRIOR APPLICATION NUMBER: 60/189199  
; PRIOR FILING DATE: 2000-03-14  
; NUMBER OF SEQ ID NOS: 12  
; SOFTWARE: PatentIn version 3.1  
; SEQ ID NO 6  
; LENGTH: 419  
; TYPE: PRT  
; ORGANISM: Homo sapiens  
US-10-182-263-6  
Query Match 90.9%; Score 179; DB 9; Length 419;  
Best Local Similarity 79.5%; Pred. No. 5.4e-22;  
Matches 35; Conservative 0; Mismatches 9; Indels 0; Gaps 0;  
QY 1 ANSFLXLRQGSIXRXCIXXICDFXKXKXIFEDVDITLAFWSKH 44  
DB 1 ANSFLXLRQGSIXRXCIXXICDFXKXKXIFEDVDITLAFWSKH 44  
RESULT 2  
US-10-182-263-3  
; Sequence 3, Application US/10182263  
; Publication No. US20030022354A1  
; GENERAL INFORMATION:  
; APPLICANT: Gerlitz, Bruce E  
; APPLICANT: Jones, Bryan E  
; APPLICANT: Grinnell, Brian W  
; TITLE OF INVENTION: PROTEIN C DERIVATIVES

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FILE REFERENCE: X-13611
; CURRENT APPLICATION NUMBER: US/10/182,263
; CURRENT FILING DATE: 2002-07-22
; PRIOR APPLICATION NUMBER: 60/181948
; PRIOR FILING DATE: 2002-02-11
; PRIOR APPLICATION NUMBER: 60/189199
; PRIOR FILING DATE: 2000-03-14
; NUMBER OF SEQ ID NOS: 12
; SOFTWARE: Patentin version 3.1
; SEQ ID NO 3
; LENGTH: 419
; TYPE: PRT
; ORGANISM: Homo sapiens
US-10-182-263-3

Query Match
Best Local Similarity 88.3%; Score 174; DB 9; Length 419;
Matches 34; Conservative 0; Mismatches 10; Indels 0; Gaps 0;

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Db 1 ANSFLELRHGSLSRECIIEICDFEAKEIFEDVDDTLAFWSKH 44

RESULT 3
US-10-182-263-4
; Sequence 4, Application US/10182263
; Publication No. US20030022354A1
; GENERAL INFORMATION:
; APPLICANT: Gerlitz, Bruce E
; APPLICANT: Jones, Bryan E
; APPLICANT: Grinnell, Brian W
; TITLE OF INVENTION: PROTEIN C DERIVATIVES
; FILE REFERENCE: X-13611
; CURRENT APPLICATION NUMBER: US/10/182,263
; CURRENT FILING DATE: 2002-07-22
; PRIOR APPLICATION NUMBER: 60/181948
; PRIOR FILING DATE: 2002-02-11
; PRIOR APPLICATION NUMBER: 60/189199
; PRIOR FILING DATE: 2000-03-14
; NUMBER OF SEQ ID NOS: 12
; SOFTWARE: Patentin version 3.1
; SEQ ID NO 4
; LENGTH: 419
; TYPE: PRT
; ORGANISM: Homo sapiens
US-10-182-263-4

Query Match
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Matches 34; Conservative 0; Mismatches 10; Indels 0; Gaps 0;

OY 1 ANSFLXXLRQGSIXRCIXXICDFFXXAKXIFEDVDDTLAFWSKH 44
Db 1 ANSFLELRHGSLSRECIIEICDFEAKEIFEDVDDTLAFWSKH 44

RESULT 4
US-10-182-263-5
; Sequence 5, Application US/10182263
; Publication No. US20030022354A1
; GENERAL INFORMATION:
; APPLICANT: Gerlitz, Bruce E
; APPLICANT: Jones, Bryan E
; APPLICANT: Grinnell, Brian W
; TITLE OF INVENTION: PROTEIN C DERIVATIVES
; FILE REFERENCE: X-13611
; CURRENT APPLICATION NUMBER: US/10/182,263
; CURRENT FILING DATE: 2002-07-22
; PRIOR APPLICATION NUMBER: 60/181948
; PRIOR FILING DATE: 2002-02-11
; PRIOR APPLICATION NUMBER: 60/189199
; PRIOR FILING DATE: 2000-03-14
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NUMBER OF SEQ ID NOS: 12
; SOFTWARE: Patentin version 3.1
; SEQ ID NO 5
; LENGTH: 419
; TYPE: PRT
; ORGANISM: Homo sapiens
US-10-182-263-5

Query Match
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Matches 34; Conservative 0; Mismatches 10; Indels 0; Gaps 0;

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Db 1 ANSFLELRHGSLSRECIIEICDFEAKEIFEDVDDTLAFWSKH 44

RESULT 5
US-10-182-263-1
; Sequence 1, Application US/10182263
; Publication No. US20030022354A1
; GENERAL INFORMATION:
; APPLICANT: Gerlitz, Bruce E
; APPLICANT: Jones, Bryan E
; APPLICANT: Grinnell, Brian W
; TITLE OF INVENTION: PROTEIN C DERIVATIVES
; FILE REFERENCE: X-13611
; CURRENT APPLICATION NUMBER: US/10/182,263
; CURRENT FILING DATE: 2002-07-22
; PRIOR APPLICATION NUMBER: 60/181948
; PRIOR FILING DATE: 2002-02-11
; PRIOR APPLICATION NUMBER: 60/189199
; PRIOR FILING DATE: 2000-03-14
; NUMBER OF SEQ ID NOS: 12
; SOFTWARE: Patentin version 3.1
; SEQ ID NO 1
; LENGTH: 419
; TYPE: PRT
; ORGANISM: Homo sapiens
US-10-182-263-1

Query Match
Best Local Similarity 81.2%; Score 160; DB 9; Length 419;
Matches 31; Conservative 2; Mismatches 11; Indels 0; Gaps 0;

OY 1 ANSFLXXLRQGSIXRCIXXICDFFXXAKXIFEDVDDTLAFWSKH 44
Db 1 ANSFLELRHGSLSRECIIEICDFEAKEIFQNVDDTLAFWSKH 44

RESULT 6
US-09-978-917A-4
; Sequence 4, Application US/09978917A
; Publication No. US20030027299A1
; GENERAL INFORMATION:
; APPLICANT: Maxygen Aps; Maxygen Holdings
; TITLE OF INVENTION: Protein C or activated protein C-like molecules
; FILE REFERENCE: 0219us310 - protein C
; CURRENT APPLICATION NUMBER: US/09/978,917A
; CURRENT FILING DATE: 2001-10-17
; NUMBER OF SEQ ID NOS: 48
; SOFTWARE: Patentin Ver. 2.1
; SEQ ID NO 4
; LENGTH: 419
; TYPE: PRT
; ORGANISM: Homo sapiens
US-09-978-917A-4

Query Match
Best Local Similarity 81.2%; Score 160; DB 9; Length 419;
Matches 31; Conservative 2; Mismatches 11; Indels 0; Gaps 0;

OY 1 ANSFLXXLRQGSIXRCIXXICDFFXXAKXIFEDVDDTLAFWSKH 44
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Db 1 ANSFLXLRHSSLERECIEECDFEAKKEIFONVDDTLAFWSKH 44

RESULT 7  
US-10-182-263-2  
Sequence 2, Application US/10182263

Publication No. US20030022354A1  
GENERAL INFORMATION:  
APPLICANT: Gerlitz, Bruce E

APPLICANT: Jones, Bryan E  
APPLICANT: Glimell, Brian W  
TITLE OF INVENTION: PROTEIN C DERIVATIVES

FILE REFERENCE: X-13611  
CURRENT APPLICATION NUMBER: US/10/182,263  
CURRENT FILING DATE: 2002-07-22

PRIOR APPLICATION NUMBER: 60/181948  
PRIOR FILING DATE: 2002-02-11  
PRIOR APPLICATION NUMBER: 60/189199

PRIOR FILING DATE: 2000-03-14  
NUMBER OF SEQ ID NOS: 12  
SOFTWARE: PatentIn version 3.1

SEQ ID NO 2  
LENGTH: 461  
TYPE: PRT

ORGANISM: Homo sapiens  
US-10-182-263-2

Query Match 81.2% Score 160; DB 9; Length 461;  
Best Local Similarity 70.5%; Pred. No. 9.5e-19;  
Matches 31; Conservative 2; Mismatches 11; Indels 0; Gaps 0;

Db 43 ANSFLXLRHSSLERECIEECDFEAKKEIFONVDDTLAFWSKH 86

RESULT 8  
US-09-978-917A-2  
Sequence 2, Application US/09978917A

Publication No. US20030027299A1  
GENERAL INFORMATION:  
APPLICANT: Maxygen Aps; Maxygen Holdings

TITLE OF INVENTION: Protein C or activated protein C-like molecules  
FILE REFERENCE: 0219u310 - protein C  
CURRENT APPLICATION NUMBER: US/09/978,917A

CURRENT FILING DATE: 2001-10-17  
NUMBER OF SEQ ID NOS: 48  
SOFTWARE: PatentIn Ver. 2.1

SEQ ID NO 2  
LENGTH: 461  
TYPE: PRT

ORGANISM: Homo sapiens  
FEATURE:  
NAME/KEY: SIGNAL

LOCATION: (1)...(42)  
FEATURE:  
NAME/KEY: CHAIN

LOCATION: (43)...(461)  
US-09-978-917A-2

Query Match 81.2% Score 160; DB 9; Length 461;  
Best Local Similarity 70.5%; Pred. No. 9.5e-19;  
Matches 31; Conservative 2; Mismatches 11; Indels 0; Gaps 0;

Db 43 ANSFLXLRHSSLERECIEECDFEAKKEIFONVDDTLAFWSKH 86

RESULT 9  
US-10-017-122-2  
Sequence 2, Application US/10017122

Publication No. US20030087244A1  
GENERAL INFORMATION:  
APPLICANT: McCarthy, Jeanette

TITLE OF INVENTION: DIAGNOSIS AND TREATMENT OF VASCULAR DISEASE  
FILE REFERENCE: MMI-007  
CURRENT APPLICATION NUMBER: US/10/017,122

CURRENT FILING DATE: 2001-12-14  
PRIOR APPLICATION NUMBER: 60/337,487  
PRIOR FILING DATE: 2001-10-09

NUMBER OF SEQ ID NOS: 4  
SOFTWARE: PatentIn Ver. 2.0  
SEQ ID NO 2

LENGTH: 466  
TYPE: PRT  
ORGANISM: Homo sapiens

US-10-017-122-2

Query Match 50.3% Score 99; DB 9; Length 466;  
Best Local Similarity 48.8%; Pred. No. 1.8e-08;  
Matches 20; Conservative 4; Mismatches 17; Indels 0; Gaps 0;

Db 61 ANAFLXLRPGSLRCKXKXICDFXAKXIFEDVDDTLAFW 101

RESULT 10  
US-10-109-498-1  
Sequence 1, Application US/10109498

Publication No. US20030044908A1  
GENERAL INFORMATION:  
APPLICANT: Persson, Egon

TITLE OF INVENTION: Coagulation Factor VII Derivatives  
FILE REFERENCE: 6286,200-US  
CURRENT APPLICATION NUMBER: US/10/109,498

CURRENT FILING DATE: 2002-03-22  
PRIOR APPLICATION NUMBER: 60/281,261  
PRIOR FILING DATE: 2001-04-03

PRIOR APPLICATION NUMBER: PA 2001 00477  
PRIOR FILING DATE: 2001-03-22  
NUMBER OF SEQ ID NOS: 20

SOFTWARE: FastSeq for Windows Version 4.0  
SEQ ID NO 1  
LENGTH: 406

TYPE: PRT  
ORGANISM: Homo sapiens  
FEATURE:

NAME/KEY: VARIANT  
LOCATION: (1)...(406)  
OTHER INFORMATION: Xaa = Any Amino Acid

US-10-109-498-1

Query Match 48.7% Score 96; DB 9; Length 406;  
Best Local Similarity 70.7%; Pred. No. 5e-08;  
Matches 29; Conservative 3; Mismatches 9; Indels 0; Gaps 0;

Db 1 ANAFLXLRPGSLRCKXKXICDFXAKXIFEDVDDTLAFW 41

RESULT 11  
US-09-759-130B-313  
Sequence 313, Application US/09759130B

Publication No. US20030022279A1  
GENERAL INFORMATION:  
APPLICANT: Millennium Pharmaceuticals, Inc.

APPLICANT: McCarthy, Sean A  
APPLICANT: Fraser, Christopher C  
APPLICANT: Sharp, John D

APPLICANT: Barnes, Thomas S  
APPLICANT: Kirstl, Susan J  
APPLICANT: Mackay, Charles R

```
APPLICANT: Myers, Paul S
APPLICANT: Leiby, Kevin R
APPLICANT: Wrighton, Nicolas
APPLICANT: Goodearl, Andrew
APPLICANT: Holtzman, Douglas A
TITLE OF INVENTION: NOVEL GENES ENCODING PROTEINS HAVING
TITLE OF INVENTION: PROGNOSTIC, DIAGNOSTIC, PREVENTIVE, THERAPEUTIC, AND OTHER
FILE REFERENCE: MP100-5350NM1M
CURRENT APPLICATION NUMBER: US/09/759,130B
CURRENT FILING DATE: 2002-09-16
PRIOR APPLICATION NUMBER: US 09/479,249
PRIOR FILING DATE: 2000-01-07
PRIOR APPLICATION NUMBER: US 09/559,497
PRIOR FILING DATE: 2000-04-27
PRIOR APPLICATION NUMBER: US 09/578,063
PRIOR FILING DATE: 2000-05-24
PRIOR APPLICATION NUMBER: US 09/333,159
PRIOR FILING DATE: 1999-06-14
PRIOR APPLICATION NUMBER: US 09/596,194
PRIOR FILING DATE: 2000-07-14
PRIOR APPLICATION NUMBER: US 09/342,364
PRIOR FILING DATE: 1999-06-29
PRIOR APPLICATION NUMBER: US 09/608,452
PRIOR FILING DATE: 2000-06-30
PRIOR APPLICATION NUMBER: US 09/393,996
PRIOR FILING DATE: 1999-09-10
PRIOR APPLICATION NUMBER: US 09/602,871
PRIOR FILING DATE: 2000-06-23
PRIOR APPLICATION NUMBER: US 09/420,707
PRIOR FILING DATE: 1999-10-19
NUMBER OF SEQ ID NOS: 460
SOFTWARE: FastSeq for Windows Version 4.0
SEQ ID NO 313
LENGTH: 96
TYPE: PRT
ORGANISM: Homo sapiens
US-09-759-130B-313

Query Match      42.9%; Score 84.5; DB 9; Length 96;
Best Local Similarity 38.6%; Pred. No. 9e-07;
Matches 17; Conservative 8; Mismatches 18; Indels 1; Gaps 1;

OY 2 NSF-LXXLRGSLXRXCIXXICDFFXAKXIFEDVDTLAFMSKH 44
DB 36 NRPDLFTPGNLERECNEELCNTEAREIFVDEDKTIAFWOEY 79

RESULT 12
US-10-189-123-43
Sequence 43, Application US/10189123
Publication No. US20030082586A1
GENERAL INFORMATION:
APPLICANT: KIRST, Susan J.
APPLICANT: HOLTZMAN, Douglas A.
APPLICANT: FRASER, Christopher C.
APPLICANT: SHARP, John D.
APPLICANT: BARNES, Thomas S.
TITLE OF INVENTION: ANTIBODIES HAVING DIAGNOSTIC, PREVENTIVE, THERAPEUTIC, AND OTHER
FILE REFERENCE: 10147-1103
CURRENT APPLICATION NUMBER: US/10/189,123
CURRENT FILING DATE: 2002-07-02
PRIOR APPLICATION NUMBER: US 09/596,194
PRIOR FILING DATE: 2000-06-16
PRIOR APPLICATION NUMBER: US 09/342,364
PRIOR FILING DATE: 1999-06-29
NUMBER OF SEQ ID NOS: 100
SOFTWARE: PatentIn version 3.1
SEQ ID NO 43
LENGTH: 96
TYPE: PRT
ORGANISM: Homo sapiens
US-10-189-123-43
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Query Match      42.9%; Score 84.5; DB 9; Length 96;
Best Local Similarity 38.6%; Pred. No. 9e-07;
Matches 17; Conservative 8; Mismatches 18; Indels 1; Gaps 1;

OY 2 NSF-LXXLRGSLXRXCIXXICDFFXAKXIFEDVDTLAFMSKH 44
DB 36 NRPDLFTPGNLERECNEELCNTEAREIFVDEDKTIAFWOEY 79

RESULT 13
US-09-759-130B-312
Sequence 312, Application US/09759130B
Publication No. US20030022279A1
GENERAL INFORMATION:
APPLICANT: Millennium Pharmaceuticals, Inc.
APPLICANT: McCarthy, Sean A.
APPLICANT: Fraser, Christopher C.
APPLICANT: Sharp, John D.
APPLICANT: Barnes, Thomas S.
APPLICANT: KIRST, Susan J.
APPLICANT: Mackay, Charles R.
APPLICANT: Myers, Paul S.
APPLICANT: Leiby, Kevin R.
APPLICANT: Wrighton, Nicolas
APPLICANT: Goodearl, Andrew
APPLICANT: Holtzman, Douglas A.
TITLE OF INVENTION: NOVEL GENES ENCODING PROTEINS HAVING
TITLE OF INVENTION: PROGNOSTIC, DIAGNOSTIC, PREVENTIVE, THERAPEUTIC, AND OTHER
FILE REFERENCE: MP100-5350NM1M
CURRENT APPLICATION NUMBER: US/09/759,130B
CURRENT FILING DATE: 2002-09-16
PRIOR APPLICATION NUMBER: US 09/479,249
PRIOR FILING DATE: 2000-01-07
PRIOR APPLICATION NUMBER: US 09/559,497
PRIOR FILING DATE: 2000-04-27
PRIOR APPLICATION NUMBER: US 09/578,063
PRIOR FILING DATE: 2000-05-24
PRIOR APPLICATION NUMBER: US 09/333,159
PRIOR FILING DATE: 1999-06-14
PRIOR APPLICATION NUMBER: US 09/596,194
PRIOR FILING DATE: 2000-07-14
PRIOR APPLICATION NUMBER: US 09/342,364
PRIOR FILING DATE: 1999-06-29
PRIOR APPLICATION NUMBER: US 09/608,452
PRIOR FILING DATE: 2000-06-30
PRIOR APPLICATION NUMBER: US 09/393,996
PRIOR FILING DATE: 1999-09-10
PRIOR APPLICATION NUMBER: US 09/602,871
PRIOR FILING DATE: 2000-06-23
PRIOR APPLICATION NUMBER: US 09/420,707
PRIOR FILING DATE: 1999-10-19
NUMBER OF SEQ ID NOS: 460
SOFTWARE: FastSeq for Windows Version 4.0
SEQ ID NO 312
LENGTH: 209
TYPE: PRT
ORGANISM: Homo sapiens
US-09-759-130B-312

Query Match      42.9%; Score 84.5; DB 9; Length 209;
Best Local Similarity 38.6%; Pred. No. 2.1e-06;
Matches 17; Conservative 8; Mismatches 18; Indels 1; Gaps 1;

OY 2 NSF-LXXLRGSLXRXCIXXICDFFXAKXIFEDVDTLAFMSKH 44
DB 36 NRPDLFTPGNLERECNEELCNTEAREIFVDEDKTIAFWOEY 79

RESULT 14
US-10-189-123-42
Sequence 42, Application US/10189123
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Publication No. US20030082586A1
GENERAL INFORMATION:
APPLICANT: KIRST, Susan J.
APPLICANT: HOLTZMAN, Douglas A.
APPLICANT: FRASER, Christopher C.
APPLICANT: SHARP, John D.
APPLICANT: BARNES, Thomas S.
TITLE OF INVENTION: ANTIBODIES HAVING DIAGNOSTIC, PREVENTIVE, THERAPEUTIC, AND OTHER
FILE REFERENCE: 10147-1103
CURRENT APPLICATION NUMBER: US/10/189,123
CURRENT FILING DATE: 2002-07-02
PRIOR APPLICATION NUMBER: US 09/596,194
PRIOR FILING DATE: 2000-06-16
PRIOR APPLICATION NUMBER: US 09/342,364
PRIOR FILING DATE: 1999-06-29
NUMBER OF SEQ ID NOS: 100
SOFTWARE: Patent version 3.1
SEQ ID NO 42
LENGTH: 209
TYPE: PRT
ORGANISM: Homo sapiens
US-10-189-123-42

Query Match          42.9%; Score 84.5; DB 9; Length 209;
Best Local Similarity 38.6%; Pred. No. 2.1e-06;
Matches 17; Conservative 8; Mismatches 18; Indels 1; Gaps 1;

QY 2 NSF-LXXLRQSLXRCIXXICDFXXAKXIFEDVDTLAFWSKH 44
DB 36 NRPDLFTPGNLERECNELCNYEAREIFVEDKTIAMWOEY 79

RESULT 15
US-09-759-130B-310
Sequence 310, Application US/09759130B
GENERAL INFORMATION:
APPLICANT: Millennium Pharmaceuticals, Inc.
APPLICANT: McCarthy, Sean A.
APPLICANT: Fraser, Christopher C.
APPLICANT: Sharp, John D.
APPLICANT: Barnes, Thomas S.
APPLICANT: Kirst, Susan J.
APPLICANT: Mackay, Charles R.
APPLICANT: Myers, Paul S.
APPLICANT: Leiby, Kevin R.
APPLICANT: Wrighton, Nicolas
APPLICANT: Goodenart, Andrew
APPLICANT: Holtzman, Douglas A.
TITLE OF INVENTION: NOVEL GENES ENCODING PROTEINS HAVING
TITLE OF INVENTION: PROGNOSTIC, DIAGNOSTIC, PREVENTIVE, THERAPEUTIC, AND OTHER
FILE REFERENCE: MPI00-5350MINI
CURRENT APPLICATION NUMBER: US/09/759,130B
CURRENT FILING DATE: 2002-09-16
PRIOR APPLICATION NUMBER: US 09/479,249
PRIOR FILING DATE: 2000-01-07
PRIOR APPLICATION NUMBER: US 09/559,497
PRIOR FILING DATE: 2000-04-27
PRIOR APPLICATION NUMBER: US 09/578,063
PRIOR FILING DATE: 2000-05-24
PRIOR APPLICATION NUMBER: US 09/333,159
PRIOR FILING DATE: 1999-06-14
PRIOR APPLICATION NUMBER: US 09/596,194
PRIOR FILING DATE: 2000-07-14
PRIOR APPLICATION NUMBER: US 09/342,364
PRIOR FILING DATE: 1999-06-29
PRIOR APPLICATION NUMBER: US 09/608,452
PRIOR FILING DATE: 2000-06-30
PRIOR APPLICATION NUMBER: US 09/393,996
PRIOR FILING DATE: 1999-09-10
PRIOR APPLICATION NUMBER: US 09/602,871
PRIOR FILING DATE: 2000-06-23
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PRIOR APPLICATION NUMBER: US 09/420,707
PRIOR FILING DATE: 1999-10-19
NUMBER OF SEQ ID NOS: 460
SOFTWARE: FASTSEQ for Windows Version 4.0
SEQ ID NO 310
LENGTH: 226
TYPE: PRT
ORGANISM: Homo sapiens
US-09-759-130B-310

Query Match          42.9%; Score 84.5; DB 9; Length 226;
Best Local Similarity 38.6%; Pred. No. 2.3e-06;
Matches 17; Conservative 8; Mismatches 18; Indels 1; Gaps 1;

QY 2 NSF-LXXLRQSLXRCIXXICDFXXAKXIFEDVDTLAFWSKH 44
DB 53 NRPDLFTPGNLERECNELCNYEAREIFVEDKTIAMWOEY 96

Search completed: May 16, 2003, 10:24:33
Job time : 56 secs
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